LEXICON PHARMACEUTICALS, INC./DE Form DEF 14A March 13, 2009

UNITED STATES

SECURITIES AND EXCHANGE COMMISSION Washington, DC. 20549

SCHEDULE 14A (Rule 14A-101)

INFORMATION REQUIRED IN PROXY STATEMENT SCHEDULE 14A INFORMATION

Proxy Statement Pursuant to Section 14(a) of the Securities Exchange Act of 1934

Filed by the Registrant x Filed by a Party other than the Registrant Check the appropriate box:

- o Preliminary Proxy Statement
- x Definitive Proxy Statement
- o Definitive Additional Materials
- o Soliciting Material Pursuant to Rule 14a-11(c) or Rule 14a-12

LEXICON PHARMACEUTICALS, INC. (Name of Registrant as Specified In Its Charter)

(Name of Person(s) Filing Proxy Statement, if Other Than the Registrant)

Payment of Filing Fee (Check the appropriate box):

No fee required.

	X
()

Fee computed on table below per Exchange Act Rules 14a-6(i)(1) and 0-11.

- (1) Title of each class of securities to which transaction applies: N/A
- (2) Aggregate number of securities to which transaction applies: N/A $\,$
- (3) Per unit price or other underlying value of transaction computed pursuant to Exchange Act Rule 0-11 (set forth the amount on which the filing fee is calculated and state how it was determined): N/A
- (4) Proposed maximum aggregate value of transaction: N/A
- (5) Total fee paid: \$0

o Fee paid previously with preliminary materials: N/A

o Check box if any part of the fee is offset as provided by Exchange Act Rule 0-11(a)(2) and identify the filing for which the offsetting fee was paid previously. Identify the previous filing by registration statement number, or the

o Confidential, for Use of the Commission Only (as permitted by Rule 14a-6(e)(2))

Form or Schedule and the date of its filing.

- (1) Amount Previously Paid:(2) Form, Schedule or Registration Statement No.:
- (3) Filing Party:
- (4) Date Filed

March 13, 2009

TO OUR STOCKHOLDERS:

I am pleased to invite you to attend the 2009 annual meeting of stockholders of Lexicon Pharmaceuticals, Inc. to be held on Thursday, April 23, 2009 at 1:30 p.m., local time, at The Marriott Woodlands Waterway Hotel and Convention Center, 1601 Lake Robbins Drive, The Woodlands, Texas.

Your vote is important, regardless of the number of shares that you hold. Whether or not you plan to attend the annual meeting, I hope you will vote as soon as possible, either electronically on the Internet, by telephone or by signing and returning the enclosed proxy card. Your proxy will not be used if you are present at the annual meeting and prefer to vote in person or if you revoke your proxy.

Thank you for your ongoing support of and continued interest in Lexicon Pharmaceuticals. We look forward to seeing you at the annual meeting.

Sincerely,

/s/ Arthur T. Sands

Arthur T. Sands, M.D., Ph.D. President and Chief Executive Officer LEXICON PHARMACEUTICALS, INC. 8800 Technology Forest Place

The Woodlands, Texas 77381 (281) 863-3000

NOTICE OF ANNUAL MEETING OF STOCKHOLDERS TO BE HELD APRIL 23, 2009

TO OUR STOCKHOLDERS:

The annual meeting of stockholders of Lexicon Pharmaceuticals, Inc. will be held on Thursday, April 23, 2009 at 1:30 p.m., local time, at The Marriott Woodlands Waterway Hotel and Convention Center, 1601 Lake Robbins Drive, The Woodlands, Texas, to:

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elect three Class III directors;

- approve our Equity Incentive Plan, amending and restating our existing 2000 Equity Incentive Plan;
- approve our Non-Employee Directors' Stock Option Plan, amending and restating our existing 2000 Non-Employee Directors' Stock Option Plan;
- •ratify and approve the appointment of Ernst & Young LLP as our independent auditors for the fiscal year ending December 31, 2009; and
 - act on any other business that properly comes before the annual meeting.

You are entitled to vote at the annual meeting only if you are the record owner of shares of our common stock at the close of business on February 23, 2009.

It is important that your shares be represented at the annual meeting whether or not you plan to attend. Please cast your vote electronically on the Internet, by telephone or by signing and returning the enclosed proxy card as promptly as possible. If you are present at the annual meeting, and wish to do so, you may revoke the proxy and vote in person.

By order of the board of directors,

/s/ Jeffrey L. Wade

Jeffrey L. Wade Secretary

The Woodlands, Texas March 13, 2009

LEXICON PHARMACEUTICALS, INC. 8800 Technology Forest Place

The Woodlands, Texas 77381 (281) 863-3000

PROXY STATEMENT

FOR ANNUAL MEETING OF STOCKHOLDERS

To Be Held April 23, 2009

GENERAL INFORMATION

Purpose of this Proxy Statement

We have prepared this proxy statement to solicit proxies on behalf of our board of directors for use at our 2009 annual meeting of stockholders and any adjournment or postponement of such meeting.

Notice of Internet Availability of Proxy Materials

As permitted by rules adopted by the Securities and Exchange Commission, we are providing access to our proxy materials over the Internet. Accordingly, on or about March 13, 2009, we are mailing to our stockholders a notice containing instructions on how to access our proxy materials, including our proxy statement and annual report, and vote electronically over the Internet. The notice also provides instructions on how stockholders may request a paper copy of our proxy materials free of charge. Our proxy materials may be accessed by stockholders at any time after the date of mailing of the notice.

Time and Place of Annual Meeting

The annual meeting will be held on Thursday, April 23, 2009 at 1:30 p.m., local time, at The Marriott Woodlands Waterway Hotel and Convention Center, 1601 Lake Robbins Drive, The Woodlands, Texas.

Matters to Be Considered at the Annual Meeting

At the annual meeting, our stockholders will be asked to consider and act upon the following matters:

- the election of three Class III directors;
- a proposal to approve our Equity Incentive Plan, amending and restating our existing 2000 Equity Compensation Plan;
- a proposal to approve our Non-Employee Directors' Stock Option Plan, amending and restating our existing 2000 Non-Employee Directors' Stock Option Plan; and
- a proposal to ratify and approve the appointment of Ernst & Young LLP as our independent auditors for the fiscal year ending December 31, 2009.

Our board of directors does not intend to bring any other matters before the annual meeting and has not been informed that any other matters are to be presented by others. Our bylaws contain several requirements that must be satisfied in order for any of our stockholders to bring a proposal before one of our annual meetings, including a requirement of delivering proper advance notice to us. Stockholders are advised to review our bylaws if they intend to present a proposal at any of our annual meetings.

Record Date for Determining Entitlement to Vote

You are entitled to vote at the annual meeting if you were the record owner of shares of our common stock as of the close of business on February 23, 2009, the record date for the annual meeting established by our board of directors.

How to Vote Your Shares

You may vote in person at the annual meeting or by proxy. To ensure that your shares are represented at the annual meeting, we recommend you vote by proxy even if you plan to attend the annual meeting in person. Even if you vote by proxy, if you wish, you can revoke your proxy and vote in person at the annual meeting. If you want to vote at the annual meeting but your shares are held by an intermediary, such as a broker or bank, you will need to obtain from the intermediary either proof of your ownership of such shares as of February 23, 2009 or a proxy from such intermediary authorizing you to vote your shares at the meeting.

You may receive more than one proxy depending on how you hold your shares. If you hold your shares through an intermediary, such as a broker or bank, you may receive materials from them asking you how you want your shares to be voted at the annual meeting.

Voting by Proxy

By Internet or Telephone. You may vote electronically on the Internet or by telephone by following the instructions contained on the notice of Internet availability of our proxy materials. If you hold your shares through an intermediary, such as a broker or bank, please follow the voting instructions contained on the voting card used by the intermediary.

By Mail. If you request a paper copy of our proxy materials, you may vote by mail by completing, dating and signing the proxy card provided and mailing it in the pre-addressed envelope enclosed with the paper copy of our proxy materials.

Quorum

We must have a quorum to conduct any business at the annual meeting. This means that at least a majority of our outstanding shares eligible to vote at the annual meeting must be represented at the annual meeting, either in person or by proxy. Abstentions are counted for purposes of determining whether a quorum is present. In addition, shares of our common stock held by intermediaries that are voted for at least one matter at the annual meeting will be counted as being present for purposes of determining a quorum for all matters, even if the beneficial owner's discretion has been withheld for voting on some or all other matters (commonly referred to as a "broker non-vote").

Outstanding Shares

On the record date, we had 137,330,254 shares of our common stock outstanding. If you were the record owner of shares of our common stock on the record date, you will be entitled to one vote for each share of stock that you own on each matter that is called to vote at the annual meeting.

Vote Needed to Approve Proposals

Our Class III directors will be elected by a plurality vote. As a result, if a quorum is present at the annual meeting, the three persons receiving the greatest number of votes will be elected to serve as our Class III directors. Withholding authority to vote for a director nominee will not affect the outcome of the election of directors.

The approval of our Equity Incentive Plan and Non-Employee Directors' Stock Option Plan and the ratification and approval of the appointment of Ernst & Young LLP as our independent auditors for the year ending December 31, 2009 will require the affirmative vote of a majority of the votes cast with respect to such matters. Any other business that may properly come before the annual meeting for a vote will require the affirmative vote of a majority of the votes cast with respect to such matter unless a greater vote is required by law or our charter or bylaws. On any such matter, any abstention from voting or broker non-vote will not count as a vote for or against these proposals and will not be considered in calculating the number of votes necessary for their approval.

How Your Proxy Will Be Voted

Giving us your proxy means that you are authorizing us to vote your shares at the annual meeting in the manner you direct. You may vote for our nominees for election as Class III directors or withhold your vote for any one or more of those nominees. You may vote for or against the proposals to approve our Equity Incentive Plan and Non-Employee Directors' Stock Option Plan and ratify and approve the appointment of Ernst & Young LLP as our independent auditors for the year ending December 31, 2009 or abstain from voting on those proposals.

If you vote by proxy and do not withhold authority to vote for the election of our nominees for election as Class III directors, all of your shares will be voted for the election of those nominees. If you withhold authority to vote for one or more of our nominees for election as Class III directors, none of your shares will be voted for those nominees.

If any of our nominees for election as Class III directors become unavailable for any reason before the election, we may reduce the number of directors serving on our board of directors, or our board of directors may designate substitute nominees, as necessary. We have no reason to believe that any of our nominees for election as Class III directors will be unavailable. If our board of directors designates any substitute nominees, the persons receiving your proxy will vote your shares for such substitute(s) if they are instructed to do so by our board of directors or, in the absence of any such instructions, in accordance with their own best judgment.

If you vote by proxy but do not specify how you want your shares voted, your shares will be voted in favor of our nominees for election as Class III directors and in favor of the approval of our Equity Incentive Plan and our Non-Employee Directors' Stock Option Plan and the ratification and approval of the appointment of Ernst & Young LLP as our independent auditors for the year ending December 31, 2009.

If you vote by proxy and any additional business properly comes before the annual meeting, the persons receiving your proxy will vote your shares on those matters as instructed by our board of directors or, in the absence of any such instructions, in accordance with their own best judgment. As of the date of this proxy statement, we are not aware of any other matter to be raised at the annual meeting.

How to Revoke Your Proxy

You may revoke your proxy at any time before your shares are voted by providing our corporate secretary with either a new proxy with a later date or a written notice of your desire to revoke your proxy at the following address:

Lexicon Pharmaceuticals, Inc. 8800 Technology Forest Place The Woodlands, Texas 77381 Attention: Corporate Secretary

You may also revoke your proxy at any time prior to your shares having been voted by attending the annual meeting in person and notifying the inspector of election of your desire to revoke your proxy. Your proxy will not automatically be revoked merely because you attend the annual meeting.

Inspector of Election

Broadridge Financial Solutions, Inc. will count votes and provide a representative who will serve as an inspector of election for the annual meeting.

List of Stockholders Entitled to Vote

A list of our stockholders entitled to vote at the annual meeting will be available for inspection at the annual meeting. The stockholder list will also be available for inspection for ten days prior to the annual meeting at our corporate offices located at 8800 Technology Forest Place, The Woodlands, Texas. Any inspection of this list at our offices will need to be conducted during ordinary business hours. If you wish to conduct an inspection of the stockholder list, we request that you please contact our corporate secretary before coming to our offices.

Solicitation of Proxies and Expenses

We are asking for your proxy on behalf of our board of directors. We will bear the entire cost of preparing, printing and soliciting proxies. We will send notices of Internet availability of proxy materials and, if requested, paper copies of our proxy materials to all of our stockholders of record as of the record date and to all intermediaries, such as brokers and banks, that held any of our shares on that date on behalf of others. These intermediaries will then forward the notices and, if requested, paper copies of our proxy materials to the beneficial owners of our shares, and we will reimburse them for their reasonable out-of-pocket expenses for forwarding such materials. Our directors, officers and employees may solicit proxies by mail, in person or by telephone or other electronic communication. Our directors, officers and employees will not receive additional compensation for their solicitation efforts, but they will be reimbursed for any out-of-pocket expenses they incur.

Householding

As permitted by rules adopted by the Securities and Exchange Commission, we are delivering a single notice of Internet availability of proxy materials, annual report and proxy statement, as applicable, to any household at which two or more stockholders reside if we believe the stockholders are members of the same family, unless otherwise instructed by one or more of the stockholders. We will promptly deliver separate copies of these documents upon the written or oral request of any stockholder at a shared address to which a single copy of the documents were delivered.

If your household received a single set of any of these documents, but you would prefer to receive your own copy, or if you share an address with another stockholder and together both of you would like to receive only a single set of these documents, please follow these instructions:

- If your shares are registered in your own name, please contact our transfer agent, BNY Mellon Shareowner Services, and inform them of your request by calling them at (800) 635-9270 or writing them at 480 Washington Boulevard., Jersey City, New Jersey 07310.
- If an intermediary, such as a broker or bank, holds your shares, please contact Broadridge and inform them of your request by calling them at (800) 542-1061 or writing them at Householding Department, 51 Mercedes Way, Edgewood, New York 11717. Be sure to include your name, the name of your brokerage firm and your account number.

PROPOSAL NUMBER 1:

ELECTION OF DIRECTORS

Our board of directors, which currently has ten members, is divided or "classified" into three classes. Directors in each class are elected to hold office for a term ending on the date of the third annual meeting following the annual meeting at which they were elected. The current term of our Class III directors will expire at this annual meeting. The current terms of our Class I and Class II directors will expire at our 2010 and 2011 annual meetings of stockholders, respectively.

The board of directors has nominated and urges you to vote for the election of the individuals identified below, who have been nominated to serve as Class III directors until our 2012 annual meeting of stockholders or until their successors are duly elected and qualified. Each of these individuals is a member of our present board of directors. Your signed proxy will be voted for the nominees named below unless you specifically indicate on the proxy that you are withholding your vote.

Nominees for Class III Directors

The following individuals are nominated for election as Class III directors:

			Year First
			Became a
Name	Age	Position with the Company	Director
Arthur T. Sands, M.D.,	47	President and Chief Executive Officer	1995
Ph.D.		and Director (Class III)	
Philippe J. Amouyal	50	Director (Class III)	2007
Frank P. Palantoni	51	Director (Class III)	2004

Arthur T. Sands, M.D., Ph.D. co-founded our company and has been our president and chief executive officer and a director since September 1995. At Lexicon, Dr. Sands pioneered the development of large-scale gene knockout technology for use in drug discovery. Before founding Lexicon, Dr. Sands served as an American Cancer Society postdoctoral fellow in the Department of Human and Molecular Genetics at Baylor College of Medicine. Dr. Sands is a member of the board of directors of the Texas Institute for Genomic Medicine. He received his B.A. in economics and political science from Yale University and his M.D. and Ph.D. from Baylor College of Medicine.

Philippe J. Amouyal has been a director since August 2007 and is a managing director of The Invus Group, LLC, a position he has held since 1999. Previously, Mr. Amouyal was a vice president and director of The Boston Consulting Group, Inc. in Boston, Massachusetts, where he coordinated the global technology and electronics practice through most of the 1990s. Mr. Amouyal is a director of Weight Watchers International, Inc., as well as a number of private companies in which Invus has invested. He holds an M.S. in engineering and a DEA in management from Ecole Centrale de Paris and was a research fellow at the Center for Policy Alternatives of the Massachusetts Institute of Technology. Mr. Amouyal is a designee of Invus, L.P. pursuant to our stockholders' agreement with Invus described under the heading "Transactions with Related Persons — Arrangements with Invus."

Frank P. Palantoni has been a director since November 2004. Mr. Palantoni served as chief executive officer of Prestige Brands Holding, Inc. from April to June 2006 and as a director from January to June 2006. From 1998 to 2004, Mr. Palantoni held a variety of senior management positions with Novartis AG, most recently as president and chief executive officer, worldwide of its Gerber Products Company, Novartis Infant and Baby Division. Mr. Palantoni also served as president and chief executive officer for North American operations of Novartis Consumer Health Division from 2000 to 2001. Prior to joining Novartis, he held a series of senior management positions with The Danone Group. He holds a B.S. from Tufts University and an M.B.A. from Columbia University.

The Board of Directors recommends that stockholders vote "FOR" the foregoing nominees for election as Class III directors.

Current and Continuing Directors

The current directors of the Company are identified below:

Name	Age	Position with the Company
Arthur T. Sands, M.D., Ph.D.	47	President and Chief Executive Officer and Director
		(Class III)
Samuel L. Barker, Ph.D. (1)	66	Chairman of the Board of Directors (Class II)
Philippe J. Amouyal (2)	50	Director (Class III)
Raymond Debbane (3)	54	Director (Class I)
Robert J. Lefkowitz, M.D. (3)	65	Director (Class I)
Alan S. Nies, M.D. (2)	71	Director (Class I)
Frank P. Palantoni (1) (2)	51	Director (Class III)
Christopher J. Sobecki	50	Director (Class II)
Judith L. Swain, M.D. (3)	60	Director (Class II)
Kathleen M. Wiltsey (1)	53	Director (Class II)
(1)		Member of the Audit Committee
(0)		\mathbf{M}_{1}

(2)	Member of the Compensation Committee
(3)	Member of the Corporate Governance Committee

Information regarding the business experience of Dr. Sands, Mr. Amouyal and Mr. Palantoni is set forth above under the heading "— Nominees for Class III Directors."

Samuel L. Barker, Ph.D. has been a director since March 2000 and became chairman of our board of directors in March 2005. In March 2001, Dr. Barker co-founded Clearview Projects, Inc., a provider of partnering and transaction services to biopharmaceutical companies, and served as its president and chief executive officer from July 2003 until November 2004. Dr. Barker served in a series of leadership positions at Bristol-Myers Squibb Company until his retirement in 1999. His positions at Bristol-Myers Squibb included service as executive vice president, Worldwide Franchise Management and Strategy during 1998; president, United States Pharmaceuticals from 1992 to 1997; and president, Bristol-Myers Squibb Intercontinental Commercial Operations from 1990 to 1992. Prior to 1990, Dr. Barker held executive positions in research and development, manufacturing, finance, business development and sales and marketing at Squibb Pharmaceuticals. Dr. Barker currently serves as a director of AtheroGenics, Inc. and Cadence Pharmaceuticals, Inc. Dr. Barker received his B.S. from Henderson State College, his M.S. from the University of Arkansas and his Ph.D. from Purdue University.

Raymond Debbane has been a director since August 2007. Mr. Debbane is president and chief executive officer of The Invus Group, LLC, which he founded in New York in 1985 as the exclusive investment advisor of Benelux-based Artal Group S.A. In 1999, Artal became the controlling shareholder of Weight Watchers International, Inc., for which Mr. Debbane serves as chairman of the board of directors. He also serves as chairman or director of a number of private companies in which Invus and Artal Group S.A. have invested. Before founding The Invus Group, Mr. Debbane was a manager in the Paris office of The Boston Consulting Group, Inc., where he did consulting work for a number of major European and international companies. Mr. Debbane holds an M.B.A. from Stanford University, an M.S. in food science and technology from the University of California at Davis, and a B.S in agricultural sciences and agricultural engineering from American University of Beirut. Mr. Debbane is a designee of Invus, L.P. pursuant to our stockholders' agreement with Invus described under the heading "Transactions with Related Persons — Arrangements with Invus."

Robert J. Lefkowitz, M.D. has been a director since February 2001 and a consultant to our company since March 2003. Dr. Lefkowitz is the James B. Duke Professor of Medicine, professor of biochemistry and a Howard Hughes Medical Institute investigator at Duke University Medical Center, where he has served on the faculty since 1973. He

is a member of the National Academy of Sciences. Dr. Lefkowitz received his B.A. from Columbia University and his M.D. from Columbia University College of Physicians and Surgeons.

Alan S. Nies, M.D. has been a director since November 2003 and chairman of our medical advisory board since March 2003. From 1992 through September 2002, Dr. Nies served in a series of senior management positions at Merck & Co. Inc., most recently as senior vice president, clinical sciences from 1999 to 2002. Prior to joining Merck, Dr. Nies spent fifteen years as professor of medicine and pharmacology and head of the Division of Clinical Pharmacology at the University of Colorado Health Sciences Center. Dr. Nies holds a B.S. from Stanford University and an M.D. from Harvard Medical School. 6

Christopher J. Sobecki has been a director since August 2007 and is a managing director of The Invus Group, LLC, which he joined in 1989. Mr. Sobecki is currently a director of Weight Watchers International, Inc. and NitroMed, Inc., as well as a number of private companies in which Invus has invested. He holds a B.S. in industrial engineering from Purdue University and an M.B.A. from Harvard University. Mr. Sobecki is a designee of Invus, L.P. pursuant to our stockholders' agreement with Invus described under the heading "Transactions with Related Persons — Arrangements with Invus."

Judith L. Swain, M.D. has been a director since September 2007. Dr. Swain is the executive director of the Singapore Institute for Clinical Sciences within A*STAR, and the Lien Chow Professor of Medicine at the National University of Singapore. From 2005 to 2006, she was the dean for translational medicine at the University of California, San Diego, where she continues to maintain an appointment as an adjunct professor of medicine. Dr. Swain served as chair of the Department of Medicine at Stanford University from 1997 to 2005, and previously served on the medical faculties of the University of Pennsylvania and Duke University. Dr. Swain is currently a director of the Burroughs Wellcome Fund, a member of the Academic Research Council in Singapore and on the Scientific Advisory Board of the Doris Duke Charitable Foundation. She has served in a number of national and international leadership roles and as a director or member of the scientific advisory boards for a number of biomedical technology companies and is co-founder of Synecor, LLC. Dr. Swain received her B.S. from the University of California, Los Angeles and her M.D. from the University of California, San Diego.

Kathleen M. Wiltsey has been a director since February 2007. From 1984 through 1998, Ms. Wiltsey served in a series of senior marketing and business development positions at Amgen Inc., including as co-product development team leader and marketing director for EPOGEN ® and as vice president with responsibility for Amgen's product licensing function. From May to October 2006, Ms. Wiltsey served the X Prize Foundation as executive director for the development and launch of the Archon X PRIZE for Genomics, a global technology competition to dramatically reduce the cost of sequencing human genomes and accelerate personalized medicine. Ms. Wiltsey has served in a variety of business and corporate development advisory roles for numerous biotechnology companies and is currently a director of Sequenom, Inc. and past president of the board of The Associates of the California Institute of Technology. She holds a B.S. from the Colorado School of Mines and an M.B.A. from Harvard University.

PROPOSAL NUMBER 2:

APPROVAL OF EQUITY INCENTIVE PLAN

We use stock option awards as a part of our overall compensation program in order to align the long-term interests of our employees with those of our stockholders. These awards are made principally under our 2000 Equity Incentive Plan, the purpose of which is to secure and retain the services of employees, directors and consultants, and to provide them with incentives to exert maximum efforts for the company's success, by giving them the opportunity through the granting of stock awards to benefit from increases in the value of our common stock.

The 2000 Equity Incentive Plan will terminate in accordance with its terms on February 2, 2010. As a result, we are asking that stockholders ratify and approve an amendment and restatement of the 2000 Equity Incentive Plan that, among other things, (1) extends the term of the plan until the day before the tenth (10th) anniversary of the date the amended and restated plan is adopted by our board of directors or approved by our stockholders, whichever is earlier, (2) eliminates the "evergreen share reserve" provisions of the existing plan under which the number of reserved shares increases on an annual basis in accordance with a pre-set formula (subject to the ability of our board of directors to provide for lesser increases and a current overall cap of 30,000,000 shares), and instead reserves a fixed number of 35,000,000 shares, (3) expands the types of potential awards under the plan to include phantom stock awards and stock appreciation rights, with the objective of providing our board of directors greater flexibility, if desired, in its choice of future equity incentive awards, and (4) renames the plan the Equity Incentive Plan.

The terms of the Equity Incentive Plan are summarized below and the complete text of the plan is set forth in Appendix A to this proxy statement.

The Board of Directors recommends that stockholders vote "FOR" approval of the Equity Incentive Plan, amending and restating the existing 2000 Equity Incentive Plan.

Administration of the Plan

The Equity Incentive Plan is administered by our board of directors, or a committee appointed by the board, which determines recipients and types of options and other awards to be granted, including number of shares under the option or other award and when options may be exercised. The compensation committee of the board of directors presently administers the plan.

Awards under the Plan

The Equity Incentive Plan permits the following types of awards:

•	incentive stock options;
•	nonstatutory stock options;
•	stock bonus awards;
•	restricted stock awards;
•	phantom stock awards; and
•	stock appreciation rights.

Stock options and other awards granted under the plan are evidenced by agreements that specify the terms and conditions under which they are granted. All stock options and other awards granted under the plan are subject to the terms and conditions contained in the applicable agreement and the plan.

Eligibility

Awards other than incentive stock options may be granted to employees, directors and consultants. Incentive stock options may be granted only to employees. As of March 2, 2009, approximately 400 persons were eligible to participate in the plan, including approximately 347 employees, nine non-employee directors and 44 consultants.

Shares Subject to the Plan

The total number of shares of common stock that may be issued pursuant to stock awards under the Equity Incentive Plan shall not exceed in the aggregate 35,000,000 shares. No more than 3,500,000 shares may be issued pursuant to awards other than stock options and stock appreciation rights.

If any award expires, lapses, or is terminated or forfeited for any reason, the shares subject to that award will continue to be available for the grant of awards under the plan, provided that shares that are not delivered to the holder of an award because (1) the right to receive such shares is surrendered in a "net exercise" of an option or (2) such shares are withheld in satisfaction of the withholding of taxes incurred in connection with the exercise of an option or stock appreciation right or the issuance of shares under a stock bonus award, restricted stock award or phantom stock award, the surrendered or withheld shares will not be available for subsequent issuance under the plan. Common stock issued as or on the exercise of awards under the plan may be either authorized and unissued shares or reacquired shares.

As of March 2, 2009, there were outstanding under the plan (1) options to purchase a total of 20,009,507 shares of our common stock, (2) stock bonus awards for 534,100 shares subject to continuing vesting requirements, (3) restricted stock awards for 17,973 shares, (4) no phantom stock awards and (5) no stock appreciation rights. On a pro forma basis to reflect the increase in the number of shares reserved for issuance under the Equity Incentive Plan, 10,248,791 shares remained available for issuance of new options or awards under the plan at that date. Since the founding of our company in 1995, a total of 4,741,702 shares of our common stock have been issued under the plan and its

predecessors upon the exercise or grant of options and other awards.

Stock Options

The stock options granted under the Equity Incentive Plan are evidenced by agreements that specify the number of shares of our common stock which may be purchased at a certain specified price and contain other terms and conditions, such as vesting and termination provisions. All stock options granted under the plan are subject to the terms and conditions contained in the applicable stock option agreement and the plan.

Expiration and Termination

The term of each stock option is stated in the applicable stock option agreement. In no event, however, may a stock option be exercised more than ten years after the date the option is granted. In the case of an incentive stock option granted to a 10% stockholder, the maximum term is five years from the date the option is granted.

Option Exercise Price

The exercise price of stock options awarded under the Equity Incentive Plan is determined by the plan administrator at the time the stock option is awarded. Stock options must have an exercise price that is no less than 100% of the fair market value of our common stock on the date of grant.

The fair market value of a share of common stock on a particular date is equal to the previous day's closing sales price (or the closing bid price, if no sales were reported) of the common stock if the common stock is listed on any established stock exchange or traded on the Nasdaq Stock Market. If there is no regular public trading market for the common stock, the fair market value of the common stock is determined by the board of directors.

Consideration for Exercise of Options

The consideration to be paid for shares to be issued upon exercise of a stock option, including the method of payment, shall be determined by the administrator (and, in the case of an incentive stock option, shall be determined at the time of grant) and may consist entirely of (1) cash or (2), at the discretion of the board of directors, (a) by delivery of other common stock, (b) according to a deferred payment or other similar arrangement, (c) by means of a "net exercise" of the option, or (d) in any other form of legal consideration acceptable to the board.

Stock Bonus Awards and Restricted Stock Awards

The terms and provisions of stock bonus awards and restricted stock awards shall be as set forth in the grant instrument. A stock bonus may be awarded in consideration for past services actually rendered to the company. The purchase price for a restricted stock award shall be as the administrator shall determine, but not less than 85% of the fair market value on the date the award is granted or the date the purchase is consummated. Shares awarded under a stock bonus or restricted stock award may, but need not be subject to a repurchase or forfeiture right on behalf of the company in accordance with a vesting schedule in the event the participant's employment is terminated.

Phantom Stock Awards and Stock Appreciation Rights

The terms and provisions of phantom stock awards and stock appreciation rights shall be as set forth in the grant instrument. The price of a common stock equivalent used as the basis from which appreciation is determined for purposes of a stock appreciation right shall be as the administrator shall determine, but not less than 100% of the fair market value on the date the stock appreciation right is granted. Phantom stock awards and the exercise value of a stock appreciation right may be paid in shares of common stock, cash, a combination of common stock and cash, or other consideration, as determined by the administrator and set forth in the grant instrument.

Other Provisions

Limits on Transfer of Awards

In general, plan participants may not sell, pledge, assign, transfer or otherwise dispose of any stock options or other awards other than by will or the laws of descent or distribution and the plan participant alone may exercise his stock options or other awards during his lifetime. Awards other than incentive stock options may be transferred only if

permitted under the agreement that evidences the terms of the award.

Adjustments on Changes in Capital Structure or on Change of Control

If we effect a stock split, reverse stock split, stock dividend, redemption, combination, reclassification or other similar change affecting our capital stock, adjustments reflecting the change will be made in (1) the aggregate number of shares of common stock authorized for issuance under the plan; (2) the number of shares underlying each outstanding award; and (3) if applicable, the price per share of each award.

If a change in control transaction shall occur, the surviving or acquiring corporation shall assume all awards or provide a substitute similar award. If the surviving or acquiring corporation fails to so provide such assumption or substitution, then awards held by those participants whose employment has not been terminated will be accelerated in full and the awards will subsequently terminate if not exercised. Any other awards outstanding under the plan will terminate if not exercised (if applicable) prior to the event.

Amendment or Termination of the Plan

The board may at any time amend, alter, suspend or discontinue the Equity Incentive Plan but no amendment, alteration, suspension or discontinuation which would impair your rights under any previous grant may be made without the consent of the participant.

Term of the Plan

No stock options or other awards may be granted under the Equity Incentive Plan after the day before the tenth (10th) anniversary of the date the plan is adopted by our board of directors or approved by our stockholders, whichever is earlier.

United States Federal Income Tax Consequences

The following is a brief summary of certain of the federal income tax consequences of certain transactions under the Equity Incentive Plan based on current federal income tax laws. This summary is not intended to be exhaustive and does not describe state or local tax consequences. Additional or different federal tax consequences to the employee, director or consultant or to our company may result depending on considerations other than those described below.

Nonstatutory Stock Options

In general, optionholders will not recognize any taxable income at the time they are granted nonstatutory stock options. When an optionholder exercises a nonstatutory stock option, he or she will recognize ordinary income measured by the excess of the then fair market value of the shares over the exercise price and we will be entitled to a deduction for a corresponding amount. Different rules apply to options that have a "readily ascertainable fair market value," as that phrase is defined in regulations promulgated under Section 83 of the Internal Revenue Code of 1986.

When an optionholder sells or otherwise disposes of shares that were acquired by exercising nonstatutory stock options, any amount that the optionholder receives in excess of the sum of (1) the exercise price of the shares as of the date of exercise and (2) the amount includable in income with respect to such option, if any, such sum being the optionholder's "basis" in the shares, will, in general, be treated as a long-term or short-term capital gain, depending on the holding period of the shares. We are not entitled to any tax deduction in connection with an optionholder's sale or disposition of the shares. If an optionholder receives less than his or her basis in the shares, the loss will, in general, be treated as a long-term or short-term capital loss, depending on the holding period of the shares.

Incentive Stock Options

Optionholders will not be taxed on the grant or exercise of an incentive stock option that qualifies under Section 422 of the Internal Revenue Code, unless an alternative minimum tax liability is triggered. When an optionholder sells or otherwise makes a taxable disposition of shares that he or she acquired by exercising an incentive stock option, the optionholder will recognize a capital gain on the excess of the amount realized on disposition over the exercise price of the incentive stock option, provided that the optionholder has not disposed of the shares until at least two years after the date the option was granted and one year after the date the optionholder exercised the option. Failure to comply with these holding requirements will result in ordinary income treatment for the gain. Unless the optionholder disposes of shares received on exercise of the incentive stock option before meeting the applicable holding period requirements, we will not be entitled to a deduction with respect to the grant or exercise of the incentive stock option. 10

In the event an optionholder makes a "disposition" of the shares received on exercise of an incentive stock option before meeting the two-year or one-year holding period requirements, the gain on the disposition, to the extent of the lesser of (1) the excess of the fair market value of the shares on the date of exercise over the exercise price or (2) the excess of the amount realized on disposition over the exercise price, will be treated as ordinary income to the optionholder, and we will generally be entitled to a corresponding deduction. The balance of the gain, if any, realized on such a disposition will be treated as long-term or short-term capital gain, depending on the holding period of the shares. To the extent that an optionholder is entitled to capital gains treatment, we will not be entitled to a corresponding deduction for such gain. If the amount realized at the time of the disposition is less than the exercise price, the optionholder will not be required to treat any amount as ordinary income, provided the disposition is of a type that would give rise to a recognizable loss. In such event, the loss will be treated as a long-term or short-term capital loss depending on the holding period of the shares.

Stock Bonus Awards

In general, if an individual receives a stock bonus award, he or she will be taxed on the fair market value of the shares on the date the shares are issued. We will be generally entitled to a deduction for a corresponding amount. When a stock bonus award is subject to forfeiture restrictions, an individual will not recognize any taxable income at the time he or she is granted the award, but upon the lapse of the restrictions applicable to such award, that person will recognize ordinary income equal to the fair market value of the shares on the date the restrictions on the award lapsed, and we will be entitled to a deduction for a corresponding amount. If, upon a taxable disposition of the shares, the stockholder receives more or less than his or her basis in the shares, the gain or loss will be a long-term or short-term capital gain or loss, depending on the holding period of the shares, measured from the date that the receipt of the shares was taxable to the stockholder.

Restricted Stock Awards

In general, an individual will not recognize any taxable income at the time he or she is granted an award of restricted stock, but upon the lapse of the restrictions applicable to such award, that person will recognize ordinary income equal to the fair market value of the shares on the date the restrictions on the award lapsed less the purchase price for such shares, and we will be entitled to a deduction for a corresponding amount. If the stockholder sells or otherwise disposes of such shares in a taxable disposition, the sale or disposition will be subject to the same treatment described above for a taxable disposition of shares acquired upon an exercise of a nonstatutory stock option.

Phantom Stock Awards

In general, an individual will not recognize any taxable income at the time he or she is granted an award of phantom stock. Upon settlement of a phantom stock award, the individual will recognize ordinary income equal to the fair market value of the cash or shares actually received by the individual. We will be generally entitled to a deduction for the corresponding amount.

Stock Appreciation Rights

In general, an individual will not recognize any taxable income at the time he or she is granted stock appreciation rights. Upon exercise of a stock appreciation right, the individual will recognize ordinary income equal to the fair market value of the cash or shares received by the individual upon exercise. We will be generally entitled to a deduction for the corresponding amount.

The foregoing summary does not constitute a definitive statement of the federal income tax effects of awards granted under the Plan.

PROPOSAL NUMBER 3: APPROVAL OF NON-EMPLOYEE DIRECTORS' STOCK OPTION PLAN

We use stock option awards as a part of our overall compensation program in order to align the long-term interests of the non-employee members of our board of directors with those of our stockholders. These awards are made under our 2000 Non-Employee Directors' Stock Option Plan, the purpose of which is to secure and retain the services of non-employee directors, and to provide them with incentives to exert maximum efforts for the company's success by giving them the opportunity through the granting of nonstatutory stock options to benefit from increases in the value of our common stock.

The 2000 Non-Employee Directors' Stock Option Plan will terminate in accordance with its terms on February 2, 2010. As a result, we are asking that stockholders approve an amendment and restatement of the 2000 Non-Employee Directors' Stock Option Plan that, among other things, (a) extends the term of the plan until the day before the tenth (10th) anniversary of the date the amended and restated plan is adopted by our board of directors or approved by our stockholders, whichever is earlier, (b) eliminates the "evergreen share reserve" provisions of the existing plan under which the number of reserved shares increases on an annual basis in accordance with a pre-set formula (subject to the ability of our board of directors to provide for lesser increases), and instead reserves a fixed number of 1,200,000 shares , (c) increases the number of shares underlying the annual option grant to the non-employee chairman of our board of directors from 10,000 shares to 20,000 shares (bringing within the Non-Employee Directors' Stock Option Plan our historical practice, previously effected by making grants of options for the additional 10,000 shares to the non-employee chairman of our board of directors' stock Option Plan.

The terms of the Non-Employee Directors' Stock Option Plan are summarized below and the complete text of the plan is set forth in Appendix B to this proxy statement.

The Board of Directors recommends that stockholders vote "FOR" approval of the Non-Employee Directors' Stock Option Plan, amending and restating the existing 2000 Non-Employee Directors' Stock Option Plan.

Administration of the Plan

The plan is administered by our board of directors, which oversees the grant of options under the plan and determines the provisions of each option granted, to the extent not specified in the plan.

Awards under the Plan

The plan only permits the award of nonstatutory stock options. Stock options granted under the plan are evidenced by agreements that specify the terms and conditions under which they are granted. All stock options granted under the plan are subject to the terms and conditions contained in the applicable agreement and the plan.

Eligibility

Stock options are granted to non-employee directors under the plan. Upon the date of his or her initial election or appointment to the board of directors, each non-employee director is granted an option to purchase 30,000 shares of our common stock. Further, on the day following each annual meeting of stockholders, each person who is then a non-employee director and has been a non-employee director for at least six months is granted an option to purchase 10,000 shares of our common stock (or 20,000 shares, in the case of the non-employee chairman of our board of directors). Nine of the ten members of our board of directors are currently eligible to participate in the plan.

Shares Subject to the Plan

The total number of shares of common stock that may be issued pursuant to stock awards under the Non-Employee Directors' Stock Option Plan shall not exceed in the aggregate 1,200,000 shares.

If any stock option expires, lapses, or is terminated or forfeited for any reason, the shares subject to that stock option will continue to be available for the grant of stock options under the plan. Common stock issued on the exercise of stock options under the plan may be either authorized and unissued shares or reacquired shares.

As of March 2, 2009, there were options outstanding under the plan to purchase a total of 504,000 shares of our common stock. On a pro forma basis to reflect the increase in the number of shares reserved for issuance under the Non-Employee Directors' Stock Option Plan, 696,000 shares remained available for issuance of new options or awards under the plan at that date. To date, no shares have been issued upon the exercise of options granted under the plan.

Stock Options

The stock options granted under the plan are evidenced by agreements that specify the number of shares of our common stock which may be purchased at a certain specified price and contain other terms and conditions, such as vesting and termination provisions. All stock options granted under the plan are subject to the terms and conditions contained in the applicable stock option agreement and the plan.

Vesting and Exercisability

Stock options granted under the plan "vest," or become exercisable, as follows:

- Initial grants of options to purchase 30,000 shares of our common stock provide for vesting of 1/60th of the shares subject to the option each month after grant for five years after the date of grant.
- Annual grants of options to purchase 10,000 shares of our common stock (or 20,000 shares, in the case of the non-employee chairman of our board of directors) provide for vesting of 1/12th of the shares subject to the option each month after grant for twelve months after the date of the grant.

Expiration and Termination

Stock options granted under the plan have a term of ten years from the date of grant, subject to earlier termination upon the occurrence of certain events. In no event, however, may a stock option be exercised more than ten years after the date the option is granted.

Option Exercise Price

Stock options awarded under the plan have an exercise price of 100% of the fair market value of our common stock on the date of grant. The fair market value of a share of common stock on a particular date is equal to the previous day's closing sales price (or the closing bid price, if no sales were reported) of the common stock if the common stock is listed on any established stock exchange or traded on the Nasdaq Stock Market. If there is no regular public trading market for the common stock, the fair market value of the common stock is determined by our board of directors.

Consideration for Exercise of Options

The consideration to be paid for shares to be issued upon exercise of a stock option, including the method of payment, may be paid, to the extent permitted by law, in any combination of (1) cash, (2) delivery of other shares of our common stock, or (3) by "net exercise" of the stock option.

Other Provisions

Limits on Transfer of Stock Options

In general, non-employee directors may not sell, pledge, assign, transfer or otherwise dispose of any stock options other than (i) by will or the laws of descent or distribution, (ii) in certain circumstances, by instrument to an inter vivos or testamentary trust and (iii) by gift to a member of such non-employee director's immediate family.

Adjustments on Changes in Capital Structure or on Change of Control

If we effect a stock split, reverse stock split, stock dividend, redemption, combination, reclassification or other similar change affecting our capital stock, adjustments reflecting the change will be made in (1) the aggregate number of shares of common stock authorized for issuance under the plan; (2) the number of shares underlying each outstanding stock option; and (3) the exercise price per share subject to each outstanding stock option.

If a change in control transaction shall occur, the surviving or acquiring corporation shall assume all stock options or provide or substitute similar stock options. If the surviving or acquiring corporation refuses to so provide such assumption or substitution, then the vesting of stock options granted under the plan will be accelerated in full and the stock options will subsequently terminate if not exercised.

If a change in control transaction shall occur and the surviving or acquiring corporation assumes the stock options granted under the plan, but any non-employee director is not elected or appointed to the board of directors of the surviving or acquiring corporation, then the vesting of that non-employee director's stock options shall be accelerated by 18 months.

Amendment or Termination of the Plan

The board of directors may at any time amend, alter, suspend or discontinue the plan but no amendment, alteration, suspension or discontinuation which would impair the rights of a non-employee director under any previous grant may be made without such non-employee director's consent.

Term of the Plan

No stock options may be granted under the plan after the day before the tenth (10th) anniversary of the date the plan is adopted by the board of directors or approved by our stockholders, whichever is earlier.

United States Federal Income Tax Consequences

The following is a brief summary of certain of the federal income tax consequences of certain transactions under the plan based on current federal income tax laws. This summary is not intended to be exhaustive and does not describe state or local tax consequences. Additional or different federal tax consequences to the non-employee director or to us may result depending on considerations other than those described below.

In general, optionholders will not recognize any taxable income at the time they are granted a nonstatutory stock option. When an optionholder exercises a nonstatutory stock option, he or she will recognize ordinary income measured by the excess of the then fair market value of the shares over the exercise price and we will be entitled to a deduction for a corresponding amount. Different rules apply to options that have a "readily ascertainable fair market value," as that phrase is defined in regulations promulgated under Section 83 of the Internal Revenue Code of 1986.

When an optionholder sells or otherwise disposes of shares that were acquired by exercising a nonstatutory stock option, any amount the optionholder receives in excess of the sum of (1) the exercise price of the shares as of the date of exercise and (2) the amount includable in income with respect to such option, if any, such sum being the optionholder's "basis" in the shares, will, in general, be treated as a long term or short term capital gain, depending on the holding period of the shares. We are not entitled to any tax deduction in connection with an optionholder's sale or disposition of the shares. If an optionholder receives less than his or her basis in the shares, the loss will, in general, be treated as a long term or short term capital loss, depending on the holding period of the shares.

The foregoing summary does not constitute a definitive statement of the federal income tax effects of stock options granted under the plan.

PROPOSAL NUMBER 4: RATIFICATION AND APPROVAL OF INDEPENDENT AUDITORS

The board of directors has appointed the firm of Ernst & Young LLP as our independent auditors to make an examination of our accounts for the fiscal year ending December 31, 2009, subject to ratification by our stockholders. Representatives of Ernst & Young LLP, are expected to be present at the annual meeting, will have an opportunity to make a statement if they desire to do so and are expected to be available to respond to appropriate questions.

The Board of Directors recommends that stockholders vote "FOR" ratification and approval of the appointment of Ernst & Young LLP as our independent auditors for the fiscal year ending December 31, 2009.

Compensation of Independent Auditors

The following table presents the estimated aggregate fees billed and to be billed by Ernst & Young LLP for services performed during our last two fiscal years.

	Years Ended December 31,		
	2008		2007
Audit fees(1)	\$ 336,500	\$	446,000
Audit-related fees(2)	24,000		22,000
Tax fees			
All other fees			
	\$ 360,500	\$	468,000

(1) "Audit fees" include professional services rendered for (i) the audit of our internal control over financial reporting, as required by the Sarbanes-Oxley Act of 2002, for the fiscal years ended December 31, 2007 and 2008, (ii) the audit of our annual financial statements for the fiscal years ended December 31, 2007 and 2008, (iii) the reviews of the financial statements included in our quarterly reports on Form 10-Q for such years and (iv) the issuance of consents and other matters relating to registration statements filed by us.

(2) "Audit-related fees" include assurance or related services reasonably related to our audit for the fiscal years ended December 31, 2007 and 2008. These fees related to the audit of the financial statements of our 401(k) plan and consultation concerning financial accounting and reporting standards.

The audit committee reviewed and approved all the fees described above. As part of its duties, the audit committee has determined that the provision by Ernst & Young LLP of the non-audit services described above is compatible with maintaining the auditors' independence.

Audit Committee Pre-Approval Policies and Procedures

The audit committee has adopted policies and procedures requiring the pre-approval of all audit and non-audit services rendered by our independent auditors, either as part of the audit committee's approval of the scope of the engagement of the independent auditors or on a case-by-case basis before the independent auditors are engaged to provide each service. The audit committee's pre-approval authority may be delegated to one or more of its members, but any pre-approval decision must be reported to the full audit committee at its next regularly scheduled meeting.

Audit Committee Report

The role of the audit committee is to assist the board of directors in its oversight of our financial reporting process. The audit committee reviews our internal accounting procedures and consults with, and reviews the services provided by, our independent auditors.

The management of our company is responsible for the preparation, presentation and integrity of our financial statements, our accounting and financial reporting principles and internal controls and procedures designed to assure compliance with the accounting standards and applicable laws and regulations. Our independent auditors are responsible for auditing our financial statements and expressing an opinion as to their conformity with generally accepted accounting principles.

In the performance of its oversight function, the audit committee has reviewed and discussed the audited financial statements with management. The committee has also discussed with our independent auditors the matters required to be discussed by Statement on Auditing Standards No. 61, Communication with Audit Committees, as currently in effect. Finally, the committee has received the written disclosures and the letter from our independent auditors required by the applicable requirements of the Public Company Accounting Oversight Board regarding our independent auditors' communications with the audit committee concerning independence, as currently in effect, and has discussed with our independent auditors their independence.

Based upon the review and discussions described in this report, and subject to the limitations on the role and responsibilities of the audit committee referred to in the audit committee charter, the audit committee recommended to the board of directors that the audited financial statements be included in our annual report on Form 10-K for the year ended December 31, 2008.

Audit Committee

Samuel L. Barker, Ph.D. (Chairman) Frank P. Palantoni Kathleen M. Wiltsey

The foregoing audit committee report shall not be deemed "filed" for purposes of Section 18 of the Securities Exchange Act of 1934, or otherwise subject to the liabilities of that section, nor shall it be deemed incorporated by reference by any general statement incorporating by reference this proxy statement into any filing under the Securities Act of 1933 or under the Securities Exchange Act of 1934, except to the extent that we specifically incorporate this compensation committee report by reference, and shall not otherwise be deemed filed under such acts.

STOCK OWNERSHIP OF CERTAIN BENEFICIAL OWNERS AND MANAGEMENT

The following table presents information regarding the beneficial ownership of our common stock as of February 23, 2009 by:

• each of the individuals listed in "Executive and Director Compensation — Summary Compensation Table";

each of our directors;

- •each person, or group of affiliated persons, who is known by us to own beneficially five percent or more of our common stock; and
 - •

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all current directors and executive officers as a group.

Beneficial ownership is determined in accordance with the rules of the Securities and Exchange Commission computing the number of shares beneficially owned by a person and the percentage ownership of that person. Shares of common stock under options held by that person that are currently exercisable or exercisable within 60 days of February 23, 2009 are considered outstanding. These shares, however, are not considered outstanding when computing the percentage ownership of each other person.

Except as indicated in the footnotes to this table and pursuant to state community property laws, each stockholder named in the table has sole voting and investment power for the shares shown as beneficially owned by them. Percentage of ownership is based on 137,330,254 shares of common stock outstanding on February 23, 2009. Unless otherwise indicated in the footnotes, the address of each of the individuals named below is: c/o Lexicon Pharmaceuticals, Inc., 8800 Technology Forest Place, The Woodlands, Texas 77381.

	Number of Shares Beneficially Owned	Beneficial Ownership Shares Issuable Pursuant to Options Exercisable within 60 Days of February 23, 2009	Percentage Ownership
Invus, L.P., Invus Public Equities, L.P. and related	55 205 1 46		10.29
parties (1)	55,385,146		40.3%
Royce & Associates, LLC (2)	9,205,777		6.7%
Arthur T. Sands, M.D., Ph.D. (3)	1,663,562	1,912,813	2.6%
Alan J. Main, Ph.D.	44,800	681,982	*
Jeffrey L. Wade, J.D.	47,800	642,367	*
Brian P. Zambrowicz, Ph.D.	156,800	890,344	*
James F. Tessmer	25,800	98,486	*
Julia P. Gregory (4)	51,983	982,112	*
Samuel L. Barker, Ph.D. (5)	47,000	134,000	*
Philippe J. Amouyal		19,500	*
Raymond Debbane (6)	55,385,146	19,500	40.3%
Robert J. Lefkowitz, M.D.		88,000	*
Alan S. Nies, M.D.	5,000	78,500	*
Frank P. Palantoni		56,500	*

1,000	19,500	*
	19,500	*
	22,500	*
57,414,408	4,851,421	43.8%
	,	19,500 22,500

* Represents beneficial ownership of less than 1 percent.

(1)Based upon a Schedule 13D filed with the SEC on June 27, 2007, and amended on August 24 and August 29, 2007, and certain representations made to us by Invus, L.P. Reflects the beneficial ownership of (a) 51,494,038 shares of our common stock by Invus, L.P., Invus Advisors, L.L.C., Ulys, L.L.C. and Raymond Debbane, each of which may be deemed to have sole voting and investment power with respect to such shares, and (b) 3,891,108 shares of our common stock by Invus Public Equities, L.P., Invus Public Equities Advisors, LLC, Ulys, L.L.C. and Mr. Debbane, each of which may be deemed to have sole voting restrictions pursuant to our stockholders' agreement with Invus, L.P. described under the heading "Transactions with Related Persons — Arrangements with Invus." The address for Invus, L.P., Invus Advisors, L.L.C., Invus Public Equities, L.P., Invus Public Equities Advisors, LLC, Ulys, L.L.C. is 750 Lexington Avenue, 30th Floor, New York, New York 10022.

- (2)Based upon a Schedule 13G filed with the SEC on January 26, 2009, reflecting the beneficial ownership of our common stock by Royce & Associates, LLC. The address for Royce & Associates, LLC is 1414 Avenue of the Americas, New York, New York 10019.
- (3) The number of shares beneficially owned by Dr. Sands includes 60,000 shares held in the name of minor children and 817,500 shares owned by Sands Associates LP. The general partners of Sands Associates LP are ATS Associates, L.L.C., owned by Dr. Sands, and MES Associates, L.L.C., owned by Dr. Sands' wife.
- (4)Based on certain representations made to us by Ms. Gregory. The number of shares beneficially owned by Ms. Gregory includes 4,847 shares held in trusts for the benefit of her children, of which she serves as trustee.
- (5) The number of shares beneficially owned by Dr. Barker includes 35,000 shares held in grantor retained annuity trusts for the benefit of his children, of which he serves as trustee.
- (6) Based upon a Schedule 13D filed with the SEC on June 27, 2007, and amended on August 24 and August 29, 2007, and certain representations made to us by Mr. Debbane. Mr. Debbane disclaims beneficial ownership of these shares. The address for Mr. Debbane is c/o Ulys, L.L.C., 750 Lexington Avenue, 30th Floor, New York, New York 10022.

Section 16(a) Beneficial Ownership Reporting Compliance

Section 16(a) of the Exchange Act requires our directors and executive officers, and persons who own more than 10% of our common stock, to file initial reports of ownership and reports of changes in ownership of our common stock with the Securities and Exchange Commission. Directors, executive officers and greater than 10% stockholders are required by Securities and Exchange Commission regulations to furnish us with copies of all such forms that they file.

To our knowledge, based solely on our review of the copies of such reports received by us and on written representations by certain reporting persons that no reports on Form 5 were required, we believe that during the fiscal year ended December 31, 2008, all Section 16(a) filing requirements applicable to our executive officers, directors and 10% stockholders were complied with in a timely manner.

EQUITY COMPENSATION PLAN INFORMATION

The following table presents aggregate summary information as of December 31, 2008 regarding the common stock that may be issued upon exercise of options, warrants and rights under all of our existing equity compensation plans, including our 2000 Equity Incentive Plan, 2000 Non-Employee Directors' Stock Option Plan and Coelacanth Corporation 1999 Stock Option Plan.

	(a)	(b)	(c)
	Number of	Weighted	
	securities to be	average exercise	Number of securities remaining
	issued upon exercise		available for future issuance
	of outstanding of outstanding		under equity compensation plans
	options, warrants	options, warrants	Č,
Plan Category	and rights	and rights	column (a))
Equity compensation plans approved by			
security holders (1)	16,840,404	\$ 5.1398	5,601,994 (3)(4)(5)
Equity compensation plans not approved			
by security holders (2)	57,760	2.2631	_

Total	16,898,164	\$ 5.1300	5,601,994

- (1)Consists of shares of our common stock issuable upon the exercise of options granted under our 2000 Equity Incentive Plan and 2000 Non-Employee Directors' Stock Option Plan or remaining available for issuance under those plans.
- (2)Consists of shares of our common stock issuable upon the exercise of options granted under the Coelacanth Corporation 1999 Stock Option Plan, which we assumed in connection with our July 2001 acquisition of Coelacanth Corporation, but does not include warrants to purchase 16,483 shares of common stock at a weighted average exercise price of \$11.93 per share, which we also assumed in connection with our acquisition of Coelacanth.
- (3)Includes 5,473,967 shares available for future issuance under our 2000 Equity Incentive Plan, some or all of which may be awarded as stock bonuses.
- (4)Our 2000 Equity Incentive Plan provides that on each January 1, the number of shares available for issuance under the plan will be automatically increased by the greater of (i) five percent of our outstanding shares on a fully-diluted basis or (ii) the number of shares that could be issued under awards granted under the plan during the prior year. Our board of directors may provide for a lesser increase in the number of shares available for issuance under the plan.

(5)Our 2000 Non-Employee Directors' Stock Option Plan provides that on the day following each annual meeting of stockholders, the number of shares available for issuance under the plan will be automatically increased by the greater of (i) 0.3% of our outstanding shares on a fully-diluted basis or (ii) the number of shares that could be issued under options granted under the plan during the prior year. Our board of directors may provide for a lesser increase in the number of shares available for issuance under the plan.

CORPORATE GOVERNANCE

Independence of the Board of Directors

After reviewing all relevant transactions and relationships involving each member of the board of directors (and his or her family), the board of directors has affirmatively determined that Samuel L. Barker, Ph.D., Philippe J. Amouyal, Raymond Debbane, Robert J. Lefkowitz, M.D., Alan S. Nies, M.D., Frank P. Palantoni, Christopher J. Sobecki, Judith L. Swain, M.D. and Kathleen M. Wiltsey, which members constitute a majority of the board of directors, are "independent" in accordance with the applicable listing standards of The Nasdaq Stock Market, Inc.

In making such determinations, the board of directors considered our consulting agreements with Robert J. Lefkowitz, M.D., under which Dr. Lefkowitz serves as a consultant to us on matters relating to our drug discovery and development efforts, and with Alan S. Nies, M.D., under which Dr. Nies serves as chairman of our medical advisory board.

Board Committees

Audit Committee. Our audit committee has been established in accordance with Section 3(a)(58)(A) of the Securities Exchange Act of 1934 and monitors the integrity of our financial statements, reviews our internal accounting procedures and oversees the qualifications, independence and performance of our independent auditors. The audit committee operates pursuant to a charter that was last amended and restated by the board of directors on October 26, 2005, a copy of which appears on our website at www.lexpharma.com under the caption "Investors — Corporate Governance."

The current members of our audit committee are Samuel L. Barker, Ph.D. (chair), Frank P. Palantoni and Kathleen M. Wiltsey. The board of directors, in its business judgment, has determined that Dr. Barker, Mr. Palantoni and Ms. Wiltsey are "independent" in accordance with the applicable listing standards of The Nasdaq Stock Market, Inc. The board of directors, in its business judgment, has also determined that Samuel L. Barker, Ph.D. is an "audit committee financial expert" as defined in Item 407(d)(5) of Regulation S-K.

Compensation Committee. Our compensation committee evaluates the performance of management, determines the compensation of our executive officers and reviews general policy relating to compensation and benefits of our employees. The compensation committee also administers the issuance of stock options and other awards under our 2000 Equity Incentive Plan. The compensation committee operates pursuant to a charter that was approved by the board of directors on February 11, 2004, a copy of which appears on our website at www.lexpharma.com under the caption "Investors — Corporate Governance."

The compensation committee may delegate any of its authority to subcommittees consisting of one or more compensation committee members, with all subcommittee decisions being presented to the full compensation committee at its next scheduled meeting. The compensation committee did not delegate any such authority with respect to 2008 compensation matters. The compensation committee may retain compensation consultants or other advisors to assist in its evaluation of executive compensation. Although the compensation committee has engaged consultants to advise the committee on matters relating to executive compensation in prior years, the compensation committee did not engage any consultants with respect to 2008 compensation matters.

The compensation committee meets in connection with most regularly scheduled meetings of the board of directors, and on at least two occasions after the commencement of each year specifically devoted to making compensation decisions regarding the year just ended. In preparation for such decisions, our president and chief executive officer reviews the performance of executive officers other than himself and, in consultation with the compensation committee and at its direction, makes certain recommendations to the compensation committee relating to their compensation. The compensation committee reviews such recommendations and makes changes to such recommendations as it deems appropriate. All executive compensation determinations are made by the compensation committee in the absence of management.

The current members of our compensation committee are Mr. Palantoni (chair), Philippe J. Amouyal and Alan S. Nies, M.D. The board of directors, in its business judgment, has determined that Mr. Palantoni, Mr. Amouyal and Dr. Nies are "independent" in accordance with the applicable listing standards of The Nasdaq Stock Market, Inc. In making such determinations, the board of directors considered our consulting agreement with Dr. Nies described under the heading "Corporate Governance — Independence of the Board of Directors."

Corporate Governance Committee. Our corporate governance committee identifies individuals qualified to become members of our board of directors, selects candidates or nominees for director positions to be filled by the board of directors or our stockholders and develops appropriate corporate governance principles. The corporate governance committee operates pursuant to a charter that was approved by the board of directors on February 11, 2004, a copy of which appears on our website at www.lexpharma.com under the caption "Investors — Corporate Governance."

The corporate governance committee has not established any specific minimum qualifications for membership on our board of directors. Rather, the committee will generally consider all relevant factors, which may include independence, experience, diversity, leadership qualities and strength of character. The corporate governance committee uses its available network of contacts when compiling a list of potential director candidates and may also engage outside consultants when appropriate. The committee also considers potential director candidates recommended by stockholders and other parties and all potential director candidates are evaluated based on the above criteria. Because the corporate governance committee makes no distinction in its evaluation of candidates based on whether such candidates are recommended by stockholders or other parties, no formal policy or procedure has been established for the consideration of director candidates recommended by stockholders.

Any stockholder wishing to propose a potential director candidate may submit a recommendation in writing within the time frame specified in our bylaws. All such communications should be sent to 8800 Technology Forest Place, The Woodlands, Texas 77381, Attn: Corporate Governance Committee. Submissions should include the full name of the proposed candidate and a detailed description of the candidate's qualifications, business experience and other relevant biographical information.

The current members of our corporate governance committee are Raymond Debbane (chair), Robert J. Lefkowitz, M.D. and Judith L. Swain, M.D. The board of directors, in its business judgment, has determined that Mr. Debbane, Dr. Lefkowitz and Dr. Swain are "independent" in accordance with the applicable listing standards of The Nasdaq Stock Market, Inc. In making such determinations, the board of directors considered our consulting agreement with Dr. Lefkowitz described under the heading "Corporate Governance — Independence of the Board of Directors."

Board and Committee Meetings and Attendance in 2008

The board of directors met six times in 2008 and took certain additional actions by unanimous written consent. The audit committee met seven times in 2008. The compensation committee met six times in 2008 and took certain additional actions by unanimous written consent. The corporate governance committee met two times in 2008. During 2008, none of our directors attended fewer than 75 percent of the aggregate number of meetings of the board of directors and committees during the period served.

It is our policy to encourage the members of our board of directors to attend all annual meetings of stockholders. Nine members of our board of directors attended our 2008 annual meeting of stockholders.

Code of Business Conduct and Ethics

We have adopted a code of business conduct and ethics that applies to all of our directors, officers and employees, the text of which appears on our website at www.lexpharma.com under the caption "Investors — Corporate Governance." We intend to disclose on our website the nature of any amendment to or waiver from our code of business conduct and

ethics that applies to our principal executive officer, principal financial officer, principal accounting officer or controller, or persons performing similar functions within four business days following the date of such amendment or waiver. In the case of any such waiver, including an implicit waiver, we also intend to disclose the name of the person to whom the waiver was granted and the date of the waiver. To date, we have not granted any waivers under our code of business conduct and ethics.

Corporate Governance Guidelines

We have adopted corporate governance guidelines, including, among other things, guidelines with respect to the structure of our board of directors, director selection and qualifications, and non-employee director compensation. The text of our corporate governance guidelines appears on our website at www.lexpharma.com under the caption "Investors — Corporate Governance."

Stockholder Communications with the Board of Directors

We believe that our stockholders are currently provided a reasonable means to communicate with our board of directors and individual directors. As a result, our board of directors has not established a formal process for stockholders to send communications to the board of directors or individual directors. However, the corporate governance committee will consider, from time to time, whether adoption of a formal process for such stockholder communications has become necessary or appropriate. Stockholders may send communications to the board of directors or individual directors by mail at 8800 Technology Forest Place, The Woodlands, Texas 77381, Attn: Board of Directors or any individual director.

Compensation Committee Interlocks and Insider Participation

During 2008, Frank P. Palantoni, Philippe J. Amouyal and Alan S. Nies, M.D. served as members of the compensation committee of our board of directors. Mr. Amouyal is a designee of Invus, L.P. pursuant to our stockholders' agreement with Invus described under the heading "Transactions with Related Persons — Arrangements with Invus." During 2008, none of our executive officers served as a member of the board of directors or compensation committee of another entity, one of whose executive officers served as a member of our board of directors or compensation committee.

TRANSACTIONS WITH RELATED PERSONS

Arrangements with Invus

In June 2007, we entered into a securities purchase agreement with Invus, L.P., pursuant to which Invus purchased 50,824,986 shares of our common stock for approximately \$205.4 million in August 2007. This purchase resulted in Invus' ownership of 40% of the post-transaction outstanding shares of our common stock. Pursuant to the securities purchase agreement, Invus also has the right to require us to initiate up to two pro rata rights offerings to our stockholders, which would provide all stockholders with non-transferable rights to acquire shares of our common stock, in an aggregate amount of up to \$344.5 million, less the proceeds of any "qualified offerings" that we may complete in the interim involving the sale of our common stock at prices above \$4.50 per share. Invus may exercise its right to require us to conduct the first rights offering by giving us notice within a period of 90 days beginning on November 28, 2009 (which we refer to as the first rights offering trigger date), although we and Invus may agree to change the first rights offering trigger date to as early as August 28, 2009 with the approval of the members of our board of directors who are not affiliated with Invus. Invus may exercise its right to require us to conduct the second rights offering by giving us notice within a period of 90 days beginning on the date that is 12 months after Invus' exercise of its right to require us to conduct the first rights offering or, if Invus does not exercise its right to require us to conduct the first rights offering, within a period of 90 days beginning on the first anniversary of the first rights offering trigger date. The initial investment and subsequent rights offerings, combined with any qualified offerings, were designed to achieve up to \$550 million in proceeds to us. Invus would participate in each rights offering for up to its pro rata portion of the offering, and would commit to purchase the entire portion of the offering not subscribed for by other stockholders.

Board of Directors. Concurrently with the execution of the securities purchase agreement, we entered into a stockholders' agreement with Invus under which Invus has the right to designate the greater of three members or 30% (or the percentage of all the outstanding shares of our common stock owned by Invus and its affiliates, if less than 30%) of all members of our board of directors, rounded up to the nearest whole number of directors, and pursuant to which Invus has designated Philippe J. Amouyal, Raymond Debbane and Christopher J. Sobecki. Mr. Debbane is president and chief executive officer of The Invus Group, LLC, an affiliate of Invus, and Mr. Amouyal and Mr. Sobecki are each managing directors of The Invus Group, LLC.

In the event that the number of shares of our common stock owned by Invus and its affiliates ever exceeds 50% of the total number of shares of our common stock then outstanding (not counting for such purpose any shares acquired by Invus from third parties in excess of 40% (or, if higher, its then pro rata amount) of the total number of outstanding shares of common stock, as permitted by the standstill provisions of the stockholders' agreement), from and after that time, Invus will have the right to designate a number of directors equal to the percentage of all the outstanding shares of our common stock owned by Invus and its affiliates (not counting for such purpose any shares acquired by Invus from third parties in excess of 40% (or, if higher, its then pro rata amount) of the total number of outstanding shares of common stock, as permitted by the standstill provisions of the stockholders' agreement), rounded up to the nearest whole number of directors. The directors appointed by Invus have proportionate representation on the compensation committee and corporate governance committee of our board of directors.

Invus' rights with respect to the designation of members of our board of directors and its compensation and corporate governance committees will terminate if the percentage of all the outstanding shares of our common stock owned by Invus and its affiliates falls below 10%. Invus will also have the right to terminate these provisions at any time following the date on which the percentage of all the outstanding shares of our common stock owned by Invus and its affiliates exceeds 50% (not counting for such purpose any shares acquired by Invus and its affiliates from third parties in excess of 40% (or, if higher, its then pro rata amount) of the total number of outstanding shares of our common stock, as permitted by the standstill provisions of the stockholders' agreement).

Preemptive Rights. Invus has preemptive rights under the stockholders' agreement to participate in future equity issuances by us (including any qualified offering), subject to certain exceptions, so as to maintain its then-current percentage ownership of our capital stock. Subject to certain limitations, Invus will be required to exercise its preemptive rights in advance with respect to certain marketed offerings, in which case it will be obligated to buy its pro rata share of the number of shares being offered in such marketed offering, including any overallotment (or such lesser amount specified in its exercise of such rights), so long as the sale of the shares were priced within a range within 10% above or below the market price on the date we notified Invus of the offering and we met certain other conditions.

The provisions of the stockholders' agreement relating to preemptive rights will terminate on the earlier to occur of August 28, 2017 and the date on which the percentage of all the outstanding shares of our common stock owned by Invus and its affiliates falls below 10%.

Standstill Provisions. Invus is subject to standstill provisions restricting its ability to purchase or otherwise acquire additional shares of common stock from third parties to an amount that would result in its ownership of our common stock not exceeding 49% of the total number of shares outstanding. These standstill provisions will not apply to the acquisitions of securities by way of stock splits, stock dividends, reclassifications, recapitalizations, or other distributions by us, acquisitions contemplated by the securities purchase agreement and the stockholders' agreement, including in the rights offerings and upon Invus' exercise of preemptive rights under the stockholders' agreement.

Except for acquisitions pursuant to the provisions described above, and subject to certain exceptions, Invus has agreed that it will not, and will cause its affiliates not to, without the approval of our unaffiliated board, directly or indirectly:

- solicit proxies to vote any of our voting securities or any voting securities of our subsidiaries;
- submit to our board of directors a written proposal for any merger, recapitalization, reorganization, business combination or other extraordinary transaction involving an acquisition of us or any of our subsidiaries or any of our or our subsidiaries' securities or assets by Invus and its affiliates;
- enter into discussions, negotiations, arrangements or understandings with any third party with respect to any of the foregoing; or

• request us or any of our representatives, directly or indirectly, to amend or waive any of these standstill provisions.

The standstill provisions of the stockholders' agreement will terminate on the earliest to occur of (a) August 28, 2017, (b) the date on which the percentage of all the outstanding shares of our common stock owned by Invus and its affiliates falls below 10%, (c) the date on which the percentage of all of the outstanding shares of our common stock owned by Invus and its affiliates exceeds 50% (not counting for such purpose any shares acquired by Invus from third parties in excess of 40% (or, if higher, its then pro rata amount) of the total number of outstanding shares of common stock, as permitted by the standstill provisions of the stockholders' agreement), (d) the date on which any third party makes a public proposal to acquire (by purchase, exchange, merger or otherwise) assets or business constituting 50% or more of our revenues, net income or assets or 50% of any class of our equity securities or our board of directors recommends or approves, or proposes to recommend or approve, any such transaction or (e) the date on which any third party third party acquires beneficial ownership (by purchase, exchange, merger or otherwise) of assets or business constituting 20% or more of our revenues, net income or assets or 20% of any class of our equity securities or our board of directors recommends or approves, or proposes to recommend or approve, any such transaction.

Sales to Third Parties. Subject to certain exceptions, Invus has agreed that neither it nor its affiliates will sell any shares of common stock to third parties that are not affiliated with Invus if, to Invus' knowledge, such transfer would result in any such third party (or any person or group including such third party) owning more than 14.9% of the total number of outstanding shares of our common stock.

The provisions of the stockholders' agreement relating to sales to third parties will terminate on the earliest to occur of (a) August 28, 2017, (b) the date on which the percentage of all the outstanding shares of our common stock owned by Invus and its affiliates falls below 10%, and (c) the date on which the percentage of all the outstanding shares of our common stock owned by Invus and its affiliates exceeds 50% (not counting for such purpose any shares acquired by Invus and its affiliates from third parties in excess of 40% (or, if higher, its then pro rata amount) of the total number of outstanding shares of our common stock, as permitted by the standstill provisions of the stockholders' agreement).

Voting of Shares. In any election of persons to serve on our board of directors, Invus will be obligated to vote all of the shares of common stock held by it and its affiliates in favor of the directors nominated by our board of directors, as long as we have complied with our obligation with respect to the designation of members of our board of directors described above and the individuals designated by Invus for election to our board of directors have been nominated, and, if applicable, are serving on our board of directors. With respect to all other matters submitted to a vote of the holders of our common stock, Invus will be obligated to vote any shares that it acquired from third parties in excess of 40% (or, if higher, its then pro rata amount) of the total number of outstanding shares of common stock, as permitted by the standstill provisions of the stockholders' agreement, in the same proportion as all the votes cast by other holders of our common stock, unless Invus and we (acting with the approval of the unaffiliated board) agree otherwise. Invus may vote all other shares of our common stock held by it in its sole discretion.

The provisions of the stockholders' agreement relating to voting will terminate on the earliest to occur of (a) August 28, 2017, (b) the date on which the percentage of all the outstanding shares of our common stock held by Invus and its affiliates falls below 10%, (c) the date on which the percentage of all outstanding shares of our common stock owned by Invus and its affiliates exceeds 50% (not counting for such purpose any shares acquired by Invus from third parties in excess of 40% (or, if higher, its then pro rata amount) of the total number of outstanding shares of our common stock, as permitted by the provisions of the stockholders' agreement), and (d) the termination of the standstill provisions in accordance with the stockholders' agreement.

Minority Protections. Invus is entitled to certain minority protections, including consent rights over (a) the creation or issuance of any new class or series of shares of our capital stock (or securities convertible into or exercisable for shares of our capital stock) having rights, preferences or privileges senior to or on parity with our common stock, (b) any amendment to our certificate of incorporation or bylaws, or amendment to the certificate of incorporation or bylaws of any of our subsidiaries, in a manner adversely affecting Invus' rights under the securities purchase agreement and the related agreements, (c) the repurchase, retirement, redemption or other acquisition of our or our subsidiaries' capital stock (or securities convertible into or exercisable for shares of our or our subsidiaries' capital stock (or securities convertible into or exercisable for shares of our or our subsidiaries' capital stock (or securities convertible into or exercisable for shares of our or our subsidiaries' capital stock (or securities convertible into or exercisable for shares of our or our subsidiaries' capital stock (or securities convertible into or exercisable for shares of our or our subsidiaries' capital stock (or securities convertible into or exercisable for shares of our or our subsidiaries' capital stock (or securities convertible into or exercisable for shares of our or our subsidiaries' capital stock (or securities convertible into or exercisable for shares of our or our subsidiaries' capital stock (or securities convertible into or exercisable for shares of our or our subsidiaries' capital stock (or securities convertible into or exercisable for shares of our or our subsidiaries' capital stock (or securities convertible into or exercisable for shares of our or our subsidiaries' capital stock (or securities convertible into or exercisable for shares of our or our subsidiaries' capital stock (or securities convertible) stock (or securities convertible) stock (or securities convertible) stock (or securities

stock), (d) any increase in the size of our board of directors to more than 12 members and (e) the adoption or proposed adoption of any stockholders' rights plan, "poison pill" or other similar plan or agreement, unless Invus is exempt from the provisions of such plan or agreement.

The provisions of the stockholders' agreement relating to minority protections will terminate on the earlier to occur of August 28, 2017 and the date on which Invus and its affiliates hold less than 15% of the total number of outstanding shares of our common stock.

Registration Rights. Concurrently with the execution of the securities purchase agreement, we entered into a registration rights agreement with Invus, pursuant to which Invus has certain demand and piggyback registration rights with respect to shares of our common stock acquired by Invus under the securities purchase agreement.

Related Party Transaction Policies

We have adopted written policies and procedures for the review, approval and ratification of interested transactions with related parties. Subject to certain exceptions provided in Item 404(a) of Regulation S-K, an "interested transaction" means any transaction, arrangement or relationship in which we are a participant and the amount involved will or may be expected to exceed \$120,000 in any calendar year, and in which any related party has or will have a direct or indirect material interest. A "related party" means (a) any executive officer, director, nominee for election as a director or any person beneficially owning five percent or more of our common stock and (b) any immediate family member of such parties.

All interested transactions are subject to the review and approval of our audit committee and if advance audit committee approval is not feasible, then the interested transaction will be considered for ratification at the audit committee's next regularly scheduled meeting. In determining whether to approve or ratify any interested transaction, the audit committee will consider, among other factors it may deem appropriate, whether the interested transaction is on terms no less favorable than terms generally available to an unaffiliated third party under similar circumstances and the extent of the related party's interest in the transaction. No director participates in any discussion or approval of an interested transaction for which he or she is a related party. On at least an annual basis, the audit committee reviews and assesses any ongoing interested transactions to ensure that the transaction remains appropriate.

EXECUTIVE AND DIRECTOR COMPENSATION

Compensation Discussion and Analysis

We have developed a compensation policy that is designed to attract and retain key executives responsible for our success and motivate management to enhance long-term stockholder value. The annual compensation package for executive and other officers typically consists primarily of three elements:

- a base salary, which reflects the responsibilities relating to the position and individual performance;
- •variable annual cash bonus awards tied to the achievement of specified corporate and individual goals and milestones, relative to pre-established bonus targets expressed as a percentage of base salary; and
 - long-term stock-based incentive awards, historically in the form of stock options.

We generally seek to set targeted total cash compensation, consisting of base salaries and annual cash bonus award targets, and total direct compensation, consisting of targeted total cash compensation and long-term stock-based incentive awards, at or near the median of a peer group of biopharmaceutical companies if such compensation level is justified by company performance, individual performance and prevailing financial conditions.

In determining peer group compensation, we use available survey data from several sources, relying principally on data from a comprehensive survey of the compensation practices of several hundred companies in the biopharmaceutical industry. We expand on this survey data with reviews of the publicly-disclosed compensation

practices of a group of biopharmaceutical companies selected for comparison purposes based on one or more factors, including number of employees, revenues, stage of development, and market capitalization. In general, we have included companies in this comparison group only if they have internal discovery capabilities and efforts. In 2006, this group of companies consisted of Curagen Corporation, deCODE genetics, Inc., Exelixis, Inc., Encysive Pharmaceuticals Inc., Human Genome Sciences, Inc., Incyte Corporation, Medarex, Inc., Millennium Pharmaceuticals, Inc., Myriad Genetics, Inc., PDL Biopharma, Inc., Vertex Pharmaceuticals Incorporated and ZymoGenetics, Inc. In 2007, this group of companies consisted of Arena Pharmaceuticals, Inc., Array Biopharma Inc., Curagen, deCODE, Exelixis, Human Genome Sciences, Incyte, Medarex, Neurocrine Biosciences, Inc., Vertex and ZymoGenetics. In 2008, this group of companies consisted of Affymax, Inc., Arena, Array, Exelixis, Human Genome Sciences, Incyte, Infinity Pharmaceuticals, Inc., Medarex, Rigel Pharmaceuticals, Inc. and ZymoGenetics. The peer group of biopharmaceutical companies for which we obtained survey data and the additional groups of companies listed above do not necessarily coincide with the companies comprising the Nasdag Biotechnology Index. In general, the compensation committee has employed the 50th and 60th percentile of the broad survey data, along with medians and averages reported by the selected group of biopharmaceutical companies, in evaluating base salaries and bonus targets. Although we acknowledge the inherent limitations in comparing our compensation practices with the compensation practices of these companies, we believe that these comparisons are useful and important points of reference in making compensation determinations. 24

All compensation decisions are made by our compensation committee pursuant to authority delegated by our board of directors. In making compensation determinations and reviewing comparative data, the compensation committee reviews total direct compensation in its totality, assigning dollar values to each of the elements of such compensation, including base salary, annual cash bonus award targets and long-term stock-based incentive awards. The committee generally allocates a greater percentage of total direct compensation to long-term stock-based incentive awards in acknowledgment of the unique challenges present in the biopharmaceutical industry and in order to reinforce the alignment of interests between our executive and other officers and our stockholders.

In determining the level and composition of compensation of each of our executive and other officers, we take into account various qualitative and quantitative indicators of corporate and individual performance. Among the challenges faced by us and other companies in the biopharmaceutical industry is the unique combination of the relatively long time period typically necessary to discover, develop and commercialize drugs and the historically low success rate in doing so. As a result, in evaluating the performance of management, the compensation committee takes into consideration such factors as the progress exhibited by our drug candidates in human clinical trials, the number and quality of drug candidates in clinical trials, the pipeline of potential drug candidates for which the required regulatory filings for the initiation of clinical trials may be filed, the value and scope of strategic collaborations and alliances with leading pharmaceutical companies, and the ability to otherwise finance our operations from external sources. In addition, the compensation committee recognizes performance and achievements that are more difficult to quantify, such as the successful supervision of major corporate projects and demonstrated leadership ability.

The compensation committee may also retain compensation consultants or other advisors when it deems appropriate to assist in its evaluation of executive compensation. The compensation committee did not engage any consultants with respect to 2006, 2007 or 2008 compensation matters.

Company Performance Criteria

We generally make executive compensation determinations in February of each year, taking into account company and individual performance over the preceding year, as well as prevailing financial conditions. The compensation committee typically meets to discuss considerations relating to executive compensation determinations several times in advance of the meeting in which such determinations are actually made.

In February 2007, the compensation committee made determinations regarding 2006 bonus awards and annual stock option grants and 2007 base salaries and bonus targets, taking into account the following factors in its evaluation of corporate performance in 2006:

- the submission of the required regulatory filings and initiation of Phase 1 clinical trials for our most advanced drug candidate, LX6171;
- the submission of the required regulatory filings for the initiation of clinical trials for another drug candidate, LX1031;
 - our progress relative to our objectives in advancing our other drug discovery and development programs;

• our performance relative to our objectives for our net use of cash in operations and for capital expenditures; and

•

our cash and investments at the end of 2006.

The committee's compensation determinations in February 2007 reflected its assessment that we largely achieved our objectives relating to our drug discovery and development programs, collectively representing what the committee considered to be our most important objectives, fell short of our objectives relating to net use of cash in operations and for capital expenditures, and achieved our objectives relating to cash and investments at the end of the year.

In February 2008, the compensation committee made determinations regarding 2007 bonus awards and annual stock option grants and 2008 base salaries and bonus targets, taking into account the following factors in its evaluation of corporate performance in 2007:

- •the completion of Phase 1 clinical trials and initiation of a Phase 2 clinical trial of our most advanced drug candidate, LX6171;
- the completion of a Phase 1a and initial Phase 1b clinical trial of another drug candidate, LX1031, with an additional Phase 1b clinical trial to be conducted in 2008;
- the submission of the required regulatory filings for the initiation of clinical trials for two of our other drug candidates, LX2931 and LX1032, with the initiation of a Phase 1 clinical trial of LX2931;
 - our progress relative to our objectives in advancing our other drug discovery and development programs;
- our performance relative to our objectives for our net use of cash in operations and for capital expenditures; and
- our cash and investments relative to our objectives at the end of 2007, reflecting the completion of our financing arrangement with Invus, L.P. described under the heading "Transactions with Related Persons Arrangements with Invus" and the establishment of our financing arrangement with Symphony Icon Holdings LLC and Symphony Icon, Inc. for the clinical development of LX6171, LX1031 and LX1032.

The committee's compensation determinations in February 2008 reflected its assessment that we achieved our objectives relating to our drug discovery and development programs, collectively representing what the committee considered to be our most important objectives, fell short of our objectives relating to net use of cash in operations and for capital expenditures and exceeded our objectives relating to cash and investments at the end of the year. Taking into account the balance of factors described above, it was the committee's assessment that our overall corporate objectives were achieved.

In February 2009, the compensation committee made determinations regarding 2008 bonus awards and annual stock option grants and 2009 base salaries and bonus targets, taking into account the following factors in its evaluation of corporate performance in 2008:

- the completion of a Phase 2a clinical trial of our LX6171 drug candidate in subjects with age-associated memory impairment, but without demonstrating significant effects on the parameters of attention or memory that were evaluated;
- the completion of Phase 1 clinical trials of another drug candidate, LX1031, and commencement of dosing in a Phase 2a clinical trial in patients with irritable bowel syndrome;
- the completion of Phase 1a and Phase 1b clinical trials of a third drug candidate, LX2931, with an additional clinical trial to be conducted in early 2009 to evaluate its interaction with methotrexate in patients with rheumatoid arthritis

in preparation for a potential Phase 2a clinical trial;

- the completion of Phase 1a and Phase 1b clinical trials of a fourth drug candidate, LX1032, with a Phase 2 clinical trial to be commenced in 2009 in patients with carcinoid syndrome;
- the submission of an investigational new drug application for one of our other drug candidates, LX4211, with the initiation of a Phase 1 clinical trial in early 2009;
- our progress relative to our objectives in advancing our other drug discovery and development programs; and
- our performance relative to our objectives for our net use of cash in operations, capital expenditures and year-end cash and investments.

The committee's compensation determinations in February 2009 reflected its assessment that we partially achieved our objectives relating to our drug discovery and development programs, collectively representing what the committee considered to be our most important objectives, and achieved our objectives relating to net use of cash in operations, capital expenditures and year-end cash and investments. Taking into account the balance of factors described above, it was the committee's assessment that our overall corporate objectives were partially achieved.

Base Salary

Base salary of executive and other officers is established through negotiation between the company and the officer at the time he or she is hired, and then subsequently adjusted when the officer's base compensation is subject to review or reconsideration. While we have entered into employment agreements with certain of our executive officers, these agreements provide that base salaries after the initial year will be reviewed and determined by the compensation committee. When establishing base salary levels for executive and other officers, the compensation committee, in accordance with its general compensation policy, considers numerous factors, including the responsibilities relating to the position, the qualifications of the executive and the relevant experience the individual brings to the company, strategic goals for which the executive has responsibility, and compensation levels of companies at a comparable stage of development who compete with us for business, scientific and executive talents. When considering increases to base salary levels for officers. No pre-determined weights are given to any one of these factors.

We left 2006 base salaries of each of our executive officers unchanged from the rates established in February 2005. We increased the base salaries of each of our executive officers in February 2007 and 2008. We left 2009 base salaries of each of our executive officers unchanged from the rates established in February 2008. The base salaries of our executive officers are generally competitive with those paid by our peer group companies, with most falling near the median for such peer group companies. In establishing base salaries for 2006, 2007, 2008 and 2009, we considered the competitiveness of our cash compensation arrangements for executive officers and our cash position and needs for the applicable year.

Incentive Compensation

Cash Bonus Awards

In addition to base salary, we may award variable annual cash bonus awards to executive and other officers depending on the extent to which certain predefined corporate and personal performance goals are achieved. These performance goals include those discussed generally above, as well as strategic and operational goals for the company as a whole. We typically consider the award of cash bonuses each February relating to performance for the preceding year. For each of our officers, the compensation committee establishes a bonus target, expressed as a percentage of base salary, which is used to determine the cash bonus amount, assuming that corporate and individual goals are fully achieved. The compensation committee retains broad discretion over the amount and payment of such awards.

In determining the cash bonus awards paid in February 2007 with respect to 2006 performance, the compensation committee included the relevant factors described above under "— Company Performance Criteria" in its evaluation of corporate performance. After taking into account these factors, the compensation committee determined that our objectives for the year had been largely but not fully achieved, and awarded bonuses for 2006 performance to executive officers other than our president and chief executive officer in amounts reflecting such partial achievement. In determining the bonus payable to Dr. Sands, our president and chief executive officer, the compensation committee also took into account the competitiveness of his overall level of compensation and his contributions to the company, and awarded a bonus for 2006 performance above his bonus target for the year. 27

In determining the cash bonus awards paid in February 2008 with respect to 2007 performance, the compensation committee included the relevant factors described above under "— Company Performance Criteria" in its evaluation of corporate performance. After taking into account these factors, the compensation committee determined that our objectives for the year had been achieved, and awarded a bonus for 2007 performance to Dr. Sands, our president and chief executive officer, in an amount reflecting such achievement. In determining the bonus payable to executive officers other than Dr. Sands, the compensation committee also took into account their contributions to the company, and awarded bonuses for 2007 performance above their bonus targets for the year.

The compensation committee determined not to award cash bonuses to our officers in February 2009 with respect to 2008 performance in light of prevailing economic conditions, the state of the financial markets, and a desire to conserve the company's cash and investment resources. Instead, the compensation committee approved the grant to our officers of restricted stock bonus awards under our 2000 Equity Incentive Plan, as described below under "— Stock-Based Awards."

Stock-Based Awards

All of our employees, including our executive and other officers, are eligible to receive long-term stock-based incentive awards under our 2000 Equity Incentive Plan as a means of providing such individuals with a continuing proprietary interest in our success. These grants are typically awarded each February and align the interests of our employees and our stockholders by providing significant incentives for our employees to achieve and maintain high levels of performance. Our 2000 Equity Incentive Plan enhances our ability to attract and retain the services of qualified individuals. Factors considered in determining whether and in what amounts such awards are granted to an officer include the executive's position, his or her performance and responsibilities, the amount of stock options currently held by the officer, the vesting schedules of any such options and the officer's other compensation. While we do not adhere to any firmly established formulas or schedules for the issuance of awards such as options or restricted stock, we take into account, in making award decisions, the total direct compensation objectives described above. In addition, we will generally tailor the terms of any such grant to achieve its goal as a long-term incentive award by providing for a vesting schedule encompassing several years.

In February 2007, 2008 and 2009, the compensation committee approved annual stock option grants to executive officers and other employees who satisfied eligibility requirements, including time of service. In making such grants, the compensation committee considered corporate and individual performance in the prior year, total direct compensation objectives for individual officers, and information regarding stock option grants made by other companies in the biotechnology industry.

As described above under "— Cash Bonus Awards," the compensation committee determined not to award cash bonuses to our officers in February 2009 with respect to 2008 performance in light of prevailing economic conditions, the state of the financial markets, and a desire to conserve the company's cash and investment resources. Instead, the compensation committee approved the grant to our officers of restricted stock bonus awards under our 2000 Equity Incentive Plan. The amounts of such awards were determined by the compensation committee based on its assessment of the achievement of the corporate and individual goals originally established for purposes of determining 2008 cash bonuses, including the relevant factors described above under "— Company Performance Criteria" in its evaluation of corporate performance. The awarded amounts reflected the compensation committee's determination that our corporate objectives for the year had been partially achieved. Taking into account both the background and nature of the awards and a desire to achieve retention and incentive objectives, the shares subject to the awards were made subject to vesting in two installments over the one-year period following the date of grant.

Summary Compensation Table for 2008

The following table presents summary information regarding the compensation of each of Arthur T. Sands, M.D., Ph.D., our principal executive officer, James F. Tessmer, our principal financial officer, and our three other most highly compensated executive officers for the year ended December 31, 2008. The table also includes summary information regarding the compensation of Julia P. Gregory, our principal financial officer until the termination of her employment on April 29, 2008. We have entered into employment agreements with certain of the named executive officers and a consulting agreement with Ms. Gregory, the material terms of which are described below. 28

Based on the summary compensation information provided below, "Salary" accounted for approximately 36%, 37% and 41% of the total compensation paid to the named executive officers in 2006, 2007 and 2008, respectively, and "Bonus" accounted for approximately 14%, 16% and 0% of the total compensation paid to the named executive officers for 2006, 2007 and 2008, respectively.

Name and Position	Year		Salary		Bonus		Stock vards (1)		Option wards (2)		All Other npensation (3)		Total
Arthur T. Sands,	I Cal		Salary		Donus	Av	valus (1)	А	walus (2)		(3)		Total
M.D., Ph.D.	2008	¢	557,500	¢		-\$	70,555	¢	900,284	¢	6,470	\$	1,534,809
President, Chief	2008	φ	557,500	φ	_	p	10,333	φ	900,204	φ	0,470	φ	1,554,609
Executive	2007	¢	522,875	¢	265,000	\$		_\$	908,842	\$	6,339	\$	1,703,056
Officer and Director			473,000		203,000	э \$			908,842 792,267	.թ \$	373,466 (4)		1,703,030
Officer and Director	2000	φ	475,000	φ	300,000	φ	_	_ \$	192,201	Φ	373,400 (4)	φ	1,930,733
Alan J. Main, Ph.D.	2008	¢	338,750	\$		_\$	30,569	\$	244,943	¢	6,236	\$	620,498
Executive Vice	2008	ψ	550,750	ψ		ψ	50,507	ψ	277,773	Ψ	0,230	ψ	020,470
President of	2007	\$	323,375	\$	130,000	\$	_	_\$	244,633	\$	6,090	\$	704,098
Pharmaceutical	2007	Ψ	525,515	Ψ	150,000	Ψ		Ψ	211,033	Ψ	0,070	Ψ	701,090
Research	2006	\$	312,000	\$	90,000	\$	_	_\$	220,103	\$	5,949	\$	628,052
Research	2000	Ψ	512,000	Ψ	,000	Ψ		Ψ	220,103	Ψ	5,515	Ψ	020,052
Jeffrey L. Wade, J.D.	2008	\$	338,333	\$	_	_\$	30,569	\$	315,287	\$	6,235	\$	690,424
Executive Vice		+		Ŧ		Ŧ	,	Ŧ	,	Ŧ	-,	+	
President and	2007	\$	316,500	\$	130,000	\$	-	_\$	326,967	\$	6,079	\$	779,546
General Counsel			292,000	\$	80,000	\$		_\$	290,675	\$	5,920	\$	668,595
					,				,				
Brian P. Zambrowicz,													
Ph.D.	2008	\$	363,333	\$	_	-\$	37,666	\$	479,148	\$	6,271	\$	886,418
Executive Vice													
President and	2007	\$	340,875	\$	140,000	\$	-	_\$	449,001	\$	6,114	\$	935,990
Chief Scientific													
Officer	2006	\$	312,000	\$	120,000	\$	-	_\$	362,435	\$	5,949	\$	800,384
James F. Tessmer	2008	\$	224,583	\$	_	-\$	16,445	\$	43,730	\$	6,073	\$	290,831
Vice President,													
Finance and													
Accounting													
Julia P. Gregory	2008	\$	115,417	\$	-	-\$	-	_\$	283,500	\$	350,461 (5)	\$	749,378
Former Executive													
Vice President	2007	\$	334,250	\$	120,000	\$	-	_\$	344,121	\$	6,106	\$	804,477
and Chief Financial													
Officer	2006	\$	329,000	\$	80,000	\$	-	_\$	332,168	\$	5,974	\$	747,142

(1)Reflects the dollar amount recognized for financial statement reporting purposes for the year ended December 31, 2008 in accordance with FAS 123(R) (but disregarding forfeiture estimates related to service-based vesting conditions) and, accordingly, includes amounts from restricted stock bonus awards granted in 2009, at our compensation committee's election, in lieu of 2008 cash bonuses.

(2) Reflects the dollar amount recognized for financial statement reporting purposes for the years ended December 31, 2006, 2007 and 2008 in accordance with FAS 123(R) (but disregarding forfeiture estimates related to service-based

vesting conditions) and, accordingly, includes amounts from options granted prior to 2006. See the information appearing under the heading entitled "Stock-Based Compensation" in footnote 2 to our consolidated financial statements included as part of our Annual Report on Form 10-K for the year ended December 31, 2008 for certain assumptions made in the valuation of options granted in the years ended December 31, 2008, 2007 and 2006. See the information appearing under the heading entitled "Stock-Based Compensation" in footnote 12 to our consolidated financial statements included as part of our Annual Report on Form 10-K for the year ended December 31, 2008, 2007 and 2006. See the information appearing under the heading entitled "Stock-Based Compensation" in footnote 12 to our consolidated financial statements included as part of our Annual Report on Form 10-K for the year ended December 31, 2005 for certain assumptions made in the valuation of options granted in the years ended December 31, 2005, 2004 and 2003.

(3)Includes the following amounts in respect of company matching contributions under our 401(k) plan and company-paid premiums for group term life insurance. The company-paid life insurance premiums reflect payments for group term life policies maintained for the benefit of all employees.

	Year	Ma Con	Company 401(k) Matching Contribution		pany-Paid oup Term Insurance emiums
Arthur T. Sands, M.D., Ph.D.	2008	\$	5,750	\$	720
	2007	\$	5,625	\$	714
	2006	\$	5,500	\$	681
Alan J. Main, Ph.D.	2008	\$	5,750	\$	486
	2003	\$	5,625	\$	465
	2006	\$	5,500	\$	449
	2000	Ψ	5,500	Ψ	
Jeffrey L. Wade, J.D.	2008	\$	5,750	\$	485
	2007	\$	5,625	\$	454
	2006	\$	5,500	\$	420
Brian P. Zambrowicz, Ph.D.	2008	\$	5,750	\$	521
	2007	\$	5,625	\$	489
	2006	\$	5,500	\$	449
James F. Tessmer	2008	\$	5,750	\$	323
Julia P. Gregory	2008	\$	5,388	\$	206
	2007	\$	5,625	\$	481
	2006	\$	5,500	\$	474

- (4) Includes reimbursable amounts foregone and payments made pursuant to a February 2006 agreement with Dr. Sands terminating our obligation under his employment agreement to fund a split-dollar life insurance policy for his benefit, pursuant to which we (a) agreed to forego our right under the split-dollar agreement with the trust that owns the policy to the reimbursement of \$147,828 in premiums that we previously paid for such policy prior to 2002 and (b) made a payment to Dr. Sands of \$219,457 enabling him to pay, for his own account, the premiums under the policy for 2004 and 2005 and the taxes associated with the termination of the trust's reimbursement obligations under the split-dollar agreement. We entered into the employment agreement with Dr. Sands in October 1999 and entered into the split-dollar agreement with the trust that owns the policy in October 2000.
- (5) Includes (a) \$38,534 paid to Ms. Gregory as compensation for her unused paid time off which had accrued at the time of her termination of employment, (b) \$25,000 paid to Ms. Gregory in consideration for her releasing us from any potential claims relating to her termination of employment, (c) \$233,333 in severance payments paid to Ms. Gregory pursuant to her employment agreement and (d) \$48,000 in consulting fees paid to Ms. Gregory pursuant to her consulting agreement. The amount of the payment made to Ms. Gregory as compensation for unused paid time off was calculated in accordance with our policies applicable to all employees.

Employment Agreements

In October 1999, we entered into an employment agreement with Arthur T. Sands, M.D., Ph.D., our president and chief executive officer, which was subsequently restated in February 2006. Under the agreement, Dr. Sands receives a base salary, currently \$560,000 a year, subject to adjustment, with an annual discretionary bonus based upon specific objectives to be determined by the compensation committee. The employment agreement is at-will and contains a non-competition agreement. The agreement also provides for certain severance payments upon the termination of Dr.

Sands' employment, as described below under the heading "Executive and Director Compensation — Potential Payments upon Termination or Change in Control."

In July 2001, we entered into an employment agreement with Alan J. Main, Ph.D., then our senior vice president, Lexicon Pharmaceuticals. In February 2007, Dr. Main was named executive vice president of pharmaceutical research. Under the agreement, Dr. Main receives a base salary, currently \$340,000 a year, subject to adjustment, with an annual discretionary bonus based upon specific objectives to be determined by the compensation committee. The employment agreement is at-will and contains a non-competition agreement. The agreement also provides for certain severance payments upon the termination of Dr. Main's employment, as described below under the heading "Executive and Director Compensation — Potential Payments upon Termination or Change in Control."

In December 1998, we entered into an employment agreement with Jeffrey L. Wade, J.D. to serve as our senior vice president and chief financial officer starting in January 1999. In February 2000, Mr. Wade was named executive vice president and general counsel. Under the agreement, Mr. Wade receives a base salary, currently \$340,000 a year, subject to adjustment, with an annual discretionary bonus based upon specific objectives to be determined by the compensation committee. The employment agreement is at-will and contains a non-competition agreement. The agreement also provides for certain severance payments upon the termination of Mr. Wade's employment, as described below under the heading "Executive and Director Compensation — Potential Payments upon Termination or Change in Control."

In February 2000, we entered into an employment agreement with Brian P. Zambrowicz, Ph.D., then our senior vice president of genomics. Dr. Zambrowicz was named executive vice president of research in August 2002 and executive vice president and chief scientific officer in February 2007. Under the agreement, Dr. Zambrowicz receives a base salary, currently \$365,000 a year, subject to adjustment, with an annual discretionary bonus based upon specific objectives to be determined by the compensation committee. The employment agreement is at-will and contains a non-competition agreement. The agreement also provides for certain severance payments upon the termination of Dr. Zambrowicz's employment, as described below under the heading "Executive and Director Compensation — Potential Payments upon Termination or Change in Control."

In February 2000, we entered into an employment agreement with Julia P. Gregory to serve as our executive vice president and chief financial officer starting in February 2000. Under the agreement, Ms. Gregory received a base salary with an annual discretionary bonus based upon specific objectives to be determined by the compensation committee. The employment agreement was at-will and contained a non-competition agreement. The agreement also contained certain severance provisions which were triggered by the termination of Ms. Gregory's employment in April 2008. Under the agreement, we are obligated to pay Ms. Gregory her most recent salary of \$350,000 per year for twelve months following the termination of her employment pursuant to our normal payroll procedures, plus an additional single sum payment of \$61,250, an amount equal to 50% of her target bonus for 2008.

Consulting Agreements

In April 2008, we entered into a consulting agreement with Ms. Gregory pursuant to which we agreed to pay Ms. Gregory \$6,000 per month for up to two days of her consulting services per month. The payments under the consulting agreement are in addition to and separate from the severance payments to which Ms. Gregory is entitled under her employment agreement. The consulting agreement has a term of one year, subject to earlier termination in certain circumstances.

Grants of Plan-Based Awards in 2008

The following table presents each grant of stock options in 2008 to the individuals named in the summary compensation table.

Name	Grant Date	Number of Securities Underlying Options	Pric of	rcise e Option wards	Fai	ant Date r Value Options
Arthur T. Sands, M.D.,						-
Ph.D.	2/2/2008	575,000	\$	2.07	\$	872,678
Alan J. Main, Ph.D.	2/7/2008	200,000	\$	2.07	\$	303,540
Jeffrey L. Wade, J.D.	2/7/2008	200,000	\$	2.07	\$	303,540
Brian P. Zambrowicz,						
Ph.D.	2/7/2008	375,000	\$	2.07	\$	569,138
James F. Tessmer	2/7/2008	40,000	\$	2.07	\$	60,708
Julia P. Gregory	2/7/2008	150,000	\$	2.07	\$	227,655

Each of the options in the foregoing table was granted under our 2000 Equity Incentive Plan and expires on the tenth anniversary of the grant date. Each option vests with respect to 25% of the shares underlying the option on the first anniversary of the grant date and 1/48th per month for each month of service thereafter. Each option becomes fully vested with respect to all remaining unvested shares upon a change in control of our company. In accordance with the process for determination of fair market value under the plan, the exercise price for each option is equal to the closing price of our common stock, as quoted on the Nasdaq Global Market, on the last trading day prior to the grant

date. The exercise price for each option may be paid in cash or in shares of our common stock valued at fair market value on the exercise date or through a cashless exercise procedure involving a same-day sale of the purchased shares.

Outstanding Equity Awards at December 31, 2008

The following table presents information about unexercised options that were held by each of the individuals listed in the summary compensation table as of December 31, 2008.

Underlying Unexercised Options Unexercisable Option Exercisable Option Exercisable Option Exercisable Option Exercisable Option Price Date Arthur T. Sands, M.D., Ph.D. 555,000 — \$ 2.50 2/3/2010 100,000 — \$ 14.44 2/2/2011 135,000 — \$ 3.30 2/14/2013 150,000 — \$ 7.59 2/12/2014 143,784 6.216 \$ 5.76 2/18/2015 269,222 110,778 \$ 4.00 2/1/3/2017 — \$ 575,000 \$ 2.07 2/7/2018 Alan J. Main, Ph.D. 49,001 — \$ 10.93 7/12/2014 15,000 — \$ 10.93 7/12/2014 15,000 — \$ 7.79 2/12/2014 47,028 2,072 \$ 5.76 2/18/2015 46,051 18.949 \$ 4.00 2/1/2016 45,840 54,160 \$ 3.390 2/14/2013 50,000 — \$ 1.73/2007 — \$ 2.50 1/13/2009 135,000 — \$ 14.44		Number of Securities								
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James F. Tessmer 30,000 — \$ 8.75 4/30/2011		91,680	,							
		—	375,000	\$	2.07	2/7/2018				
15,000 — \$ 9.60 2/7/2012	James F. Tessmer									
		15,000		\$	9.60	2/7/2012				

7,200		\$ 3.90	2/14/2013
7,600		\$ 7.59	2/12/2014
6,709	291	\$ 5.76	2/18/2015
4,675	1,925	\$ 4.00	2/1/2016
4,584	5,416	\$ 3.94	2/13/2017
7,292	17,708	\$ 3.56	10/23/2017
	40,000	\$ 2.07	2/7/2018

				Number of Securities									
						Underlying Une	exercised Options	O	ption	Option			
							Unexercisable	Ex	tercise	Expiration			
						Exercisable	(1)	I	Price	Date			
J u	1	i	а	Р		433,000	—	\$	2.50	2/8/2010			
Gregory	/												
						60,000	—	\$	14.44	2/2/2011			
						90,000		\$	9.38	2/19/2012			
						72,000	_	\$	3.90	2/14/2013			
						75,000	_	\$	7.59	2/12/2014			
						71,892	3,108	\$	5.76	2/18/2015			
						70,848	29,152	\$	4.00	2/1/2016			
						45,840	54,160	\$	3.94	2/13/2017			
						_	150,000	\$	2.07	2/7/2018			

(1)Each option vests with respect to 25% of the shares underlying the option on the first anniversary of the grant date and 1/48th per month for each month of service thereafter.

Potential Payments upon Termination or Change in Control

Employment Agreements

Arthur T. Sands, M.D., Ph.D. Our employment agreement with Dr. Sands provides that if we terminate his employment without "cause," if Dr. Sands terminates his employment for "good reason," or if his employment is terminated at the end of any renewal term through notice of non-renewal, we will pay Dr. Sands his then-current salary for twelve months pursuant to our normal payroll procedures, plus an additional single sum payment equal to his full target bonus for the year in which the termination occurred.

Under our employment agreement with Dr. Sands, "good reason" means any of the following, without Dr. Sands' prior written consent:

- any material diminution in Dr. Sands' base compensation, followed by Dr. Sands terminating his employment for "good reason" within 120 days after receiving notice of such diminution;
- any material diminution in Dr. Sands' authority, duties or responsibilities, followed by Dr. Sands terminating his employment for "good reason" within 120 days after receiving notice of such diminution;
- any requirement that Dr. Sands report to an officer or other employee of our company rather than reporting directly to our board of directors, followed by Dr. Sands terminating his employment for "good reason" within 120 days after receiving notice of such requirement; or
- any material breach by our company of the agreement, followed by Dr. Sands terminating his employment for "good reason" within 120 days after receiving notice of such breach.

Alan J. Main, Ph.D. Our employment agreement with Dr. Main provides that if we terminate his employment without "cause" or if Dr. Main terminates his employment for "good reason," we will pay Dr. Main his then-current salary for twelve months pursuant to our normal payroll procedures, plus an additional single sum payment equal to 50% of his target bonus for the year in which the termination occurred. If his employment is terminated at the end of any renewal term through notice of non-renewal, we will pay Dr. Main his then-current salary for six months pursuant to our normal payroll procedures.

Under our employment agreement with Dr. Main, "good reason" means any of the following, without Dr. Main's prior written consent:

• any material diminution in Dr. Main's base compensation, followed by Dr. Main terminating his employment for "good reason" within 120 days after receiving notice of such diminution;

- any material diminution in Dr. Main's authority, duties or responsibilities, followed by Dr. Main terminating his employment for "good reason" within 120 days after receiving notice of such diminution; or
- any material breach by our company of the agreement, followed by Dr. Main terminating his employment for "good reason" within 120 days after receiving notice of such breach.

Jeffrey L. Wade, J.D. Our employment agreement with Mr. Wade provides that if we terminate his employment without "cause" or if Mr. Wade terminates his employment for "good reason," we will pay Mr. Wade his then-current salary for twelve months pursuant to our normal payroll procedures, plus an additional single sum payment equal to 50% of his target bonus for the year in which the termination occurred, provided that if such termination occurs within 120 days following a reduction in his salary, the salary continuation payments shall be based on Mr. Wade's salary prior to such reduction. If his employment is terminated at the end of any renewal term through notice of non-renewal, we will pay Mr. Wade his then-current salary for six months pursuant to our normal payroll procedures.

Under our employment agreement with Mr. Wade, "good reason" means any of the following, without Mr. Wade's prior written consent:

- any material diminution in Mr. Wade's base compensation, followed by Mr. Wade terminating his employment for "good reason" within 120 days after receiving notice of such diminution;
- any material diminution in Mr. Wade's authority, duties or responsibilities, followed by Mr. Wade terminating his employment for "good reason" within 120 days after receiving notice of such diminution; or
- any material breach by our company of the agreement, followed by Mr. Wade terminating his employment for "good reason" within 120 days after receiving notice of such breach.

Brian P. Zambrowicz, Ph.D. Our employment agreement with Dr. Zambrowicz provides that if we terminate his employment without "cause" or if Dr. Zambrowicz terminates his employment for "good reason," we will pay Dr. Zambrowicz his then-current salary for twelve months pursuant to our normal payroll procedures, plus an additional single sum payment equal to 50% of his target bonus for the year in which the termination occurred. If his employment is terminated at the end of any renewal term through notice of non-renewal, we will pay Dr. Zambrowicz his then-current salary for six months pursuant to our normal payroll procedures.

Under our employment agreement with Dr. Zambrowicz, "good reason" means any of the following, without Dr. Zambrowicz's prior written consent:

- any material diminution in Dr. Zambrowicz's base compensation, followed by Dr. Zambrowicz terminating his employment for "good reason" within 120 days after receiving notice of such diminution;
- any material diminution in Dr. Zambrowicz's authority, duties or responsibilities, followed by Dr. Zambrowicz terminating his employment for "good reason" within 120 days after receiving notice of such diminution; or
- any material breach by our company of the agreement, followed by Dr. Zambrowicz terminating his employment for "good reason" within 120 days after receiving notice of such breach.

Julia P. Gregory. Our employment agreement with Ms. Gregory provided that if we terminated her employment without "cause," as occurred with respect to the termination of her employment in April 2008, we would pay Ms. Gregory her then-current salary for twelve months pursuant to our normal payroll procedures, plus an additional single sum payment equal to 50% of her target bonus for the year in which the termination occurred.

Under each of our employment agreements with the individuals named in the summary compensation table, "cause" means any of the following:

- the individual having engaged in intentional misconduct causing our material violation of any state or federal laws;
- the individual having engaged in a theft of corporate funds or corporate assets or in a material act of fraud upon us;
- an act of personal dishonesty taken by the individual that was intended to result in personal enrichment of the individual at our expense;
 - the individual's final conviction in a court of competent jurisdiction of a felony; or
- a breach by the individual during his or her employment of the conflict of interest, confidential information and non-competition covenants under the agreement, if such breach results in a material injury to our company.

Stock Option Agreements

Our stock option agreements with the individuals named in the summary compensation table provide that all remaining unvested stock options shall become fully vested upon a change in control of our company. Under the stock option agreements, a "change in control" shall have occurred upon any of the following events:

- any person other than Invus, L.P. and its affiliates becomes the beneficial owner of securities representing 35% or more of the combined voting power of our outstanding voting securities;
- Invus, L.P. and its affiliates become the beneficial owner of securities representing 50% or more of the combined voting power of our outstanding voting securities;
- the approval by our stockholders of a reorganization, merger, or consolidation pursuant to which our stockholders immediately prior to such reorganization, merger or consolidation do not, immediately thereafter, own or control more than 50% of the combined voting power of the surviving entity's outstanding voting securities in substantially the same proportions as prior to such reorganization, merger or consolidation;
 - our liquidation or dissolution or the sale of all or substantially all of our assets;
- the election by our stockholders of any person to our board of directors who has not been nominated for election by a majority of the board of directors or any duly appointed committee thereof;
- following the election or removal of directors, a majority of the board of directors consists of individuals who were not members of the board of directors two years before such election or removal, unless the election of such individuals to the board of directors has been approved in advance by directors representing a majority of the directors then in office who were directors at the beginning of the two-year period; or
- any other corporate event affecting the company deemed to be a "change in control" by the compensation committee.

The following table reflects the amounts Dr. Sands, Dr. Main, Mr. Wade, Dr. Zambrowicz and Mr. Tessmer would have been entitled to receive if the foregoing termination and change-in-control events had occurred on December 31, 2008. The table also reflects the amounts Ms. Gregory is entitled to receive as a result of the termination of her employment in April 2008. The table does not take into account any taxes that may have been payable in connection with any of those payments:

	-	ggregate Salary				Accelerated	k
Name	Cor	ntinuation	В	onus (6)	(Options (7)	
Arthur T. Sands, M.D., Ph.D.	\$	560,000 (1)	\$	280,000	\$		—
Alan J. Main, Ph.D.	\$	340,000 (2)	\$	59,500	\$		
Jeffrey L. Wade, J.D.	\$	340,000 (3)	\$	59,500	\$		_
Brian P. Zambrowicz, Ph.D.	\$	365,000 (4)	\$	73,000	\$		
James F. Tessmer	\$		\$		- \$		-
Julia P. Gregory	\$	350,000 (5)	\$	61,250	\$		

- (1)Reflects aggregate salary continuation payments due as a result of our termination of Dr. Sands' employment without "cause," Dr. Sands' termination of his employment for "good reason," or the termination of Dr. Sands' employment at the end of a renewal term through notice of non-renewal.
- (2)Reflects aggregate salary continuation payments due as a result of our termination of Dr. Main's employment without "cause" or Dr. Main's termination of his employment for "good reason." If Dr. Main's employment had been terminated at the end of a renewal term through notice of non-renewal, the aggregate salary continuation payment for Dr. Main would have been \$170,000.
- (3) Reflects aggregate salary continuation payments due as a result of our termination of Mr. Wade's employment without "cause" or Mr. Wade's termination of his employment for "good reason." If Mr. Wade's employment had been terminated at the end of a renewal term through notice of non-renewal, the aggregate salary continuation payment for Mr. Wade would have been \$170,000.
- (4)Reflects aggregate salary continuation payments due as a result of our termination of Dr. Zambrowicz's employment without "cause" or Dr. Zambrowicz's termination of his employment for "good reason." If Dr. Zambrowicz's employment had been terminated at the end of a renewal term through notice of non-renewal, the aggregate salary continuation payment for Dr. Zambrowicz would have been \$182,500.
- (5)Reflects aggregate salary continuation payments due as a result of our termination of Ms. Gregory's employment without "cause" in April 2008.
- (6) With respect to Dr. Sands, Dr. Main, Mr. Wade, Dr. Zambrowicz and Mr. Tessmer, reflects single-sum bonus payments due as a result of our termination of the named individual's employment without "cause," the named individual's termination of his or her employment for "good reason," or in the case of Dr. Sands, the termination of his employment at the end of a renewal term through notice of non-renewal. With respect to Ms. Gregory, reflects a single-sum bonus payment paid as a result of our termination of her employment without "cause" in April 2008.
- (7) With respect to Dr. Sands, Dr. Main, Mr. Wade, Dr. Zambrowicz and Mr. Tessmer, based on the closing price of our common stock on the Nasdaq Global Market on December 31, 2008 of \$1.40 per share, less the exercise price payable with respect to the stock options for which vesting would have been accelerated. The termination of Ms. Gregory's employment did not result in acceleration of vesting with respect to any of her then unvested stock options.

Director Compensation in 2008

Each non-employee member of our board of directors currently receives the following cash compensation:

• an annual retainer of \$15,000 for service on the board of directors (\$30,000 for service as non-executive chairman of the board of directors), prorated for any partial year of service;

- an annual retainer of \$2,500 for service on each committee of the board of directors of which he or she is a member (\$5,000 for service as chairman of any such committee), prorated for any partial year of service;
- a fee of \$2,500 for each meeting of the board of directors that he or she attends in person (\$500 for each telephonic meeting of the board of directors in which he or she participates); and
- a fee of \$1,000 for each committee meeting that he or she attends in person other than in connection with a meeting of the full board of directors (\$500 for each telephonic committee meeting in which he or she participates).

Arthur T. Sands, M.D., Ph.D., our president and chief executive officer, does not receive additional compensation for his service as a director. We make additional cash payments to Dr. Lefkowitz for his consulting services and to Dr. Nies for his consulting services as chairman of our medical advisory board.

Our 2000 Non-Employee Directors' Stock Option Plan provides for the grant of options to purchase shares of common stock to our non-employee directors. Non-employee directors first elected after the closing of our initial public offering in April 2000 receive an initial option to purchase 30,000 shares of common stock. In addition, all non-employee directors who have served in such capacity for six months receive an annual option to purchase 10,000 shares of common stock. All options granted under the non-employee directors' plan have an exercise price equal to the fair market value of our common stock on the date of grant.

The chairman of our board of directors receives an additional annual option under our 2000 Equity Incentive Plan to purchase 10,000 shares of common stock. All such options have an exercise price equal to the fair market value of our common stock on the date of grant.

The following table presents summary information for the year ended December 31, 2008 regarding the compensation of the non-employee members of our board of directors.

	Fee	s Earned or	Ol	ption Awards (1)	Al	l Other	
Name	Pa	id in Cash		(2) (3)	Com	pensation	Total
Samuel L. Barker, Ph.D.	\$	47,500	\$	39,029		— \$	86,529
Philippe J. Amouyal	\$	31,000	\$	24,124		— \$	55,124
Raymond Debbane	\$	31,000	\$	24,124		— \$	55,124
Robert J. Lefkowitz, M.D.	\$	28,500	\$	19,513	\$	50,000(4) \$	98,013
Alan S. Nies, M.D.	\$	31,000	\$	41,605	\$	75,000(5) \$	147,605
Frank P. Palantoni	\$	37,500	\$	52,349		— \$	89,849
Christopher J. Sobecki	\$	26,000	\$	24,124		— \$	50,124
Judith L. Swain, M.D.	\$	28,500	\$	24,549		— \$	53,049
Kathleen M. Wiltsey	\$	30,000	\$	29,608		— \$	59,608

- (1)Reflects the dollar amount recognized for financial statement reporting purposes for the year ended December 31, 2008 in accordance with FAS 123(R) (but disregarding forfeiture estimates related to service-based vesting conditions) and, accordingly, includes amounts from options granted prior to 2008. See the information appearing under the heading entitled "Stock-Based Compensation" in footnote 2 to our consolidated financial statements included as part of our Annual Report on Form 10-K for the year ended December 31, 2008 for certain assumptions made in the valuation of options granted in the years ended December 31, 2008, 2007 and 2006. See the information appearing under the heading entitled "Stock-Based Compensation" in footnote 12 to our consolidated financial statements included as part of our Annual Report on Form 10-K for the year ended December 31, 2008, 2007 and 2006. See the information appearing under the heading entitled "Stock-Based Compensation" in footnote 12 to our consolidated financial statements included as part of our Annual Report on Form 10-K for the year ended December 31, 2008, 2007 and 2006. See the information appearing under the heading entitled "Stock-Based Compensation" in footnote 12 to our consolidated financial statements included as part of our Annual Report on Form 10-K for the year ended December 31, 2005 for certain assumptions made in the valuation of options granted in the years ended December 31, 2005, 2004 and 2003.
- (2) The non-employee members of our board of directors who held such position on December 31, 2008 held the following aggregate number of unexercised options as of such date:

	Number of Securities Underlying Unexercised
Name	Options
Samuel L. Barker, Ph.D.	134,000
Philippe J. Amouyal	40,000
Raymond Debbane	40,000
Robert J. Lefkowitz, M.D.	88,000
Alan S. Nies, M.D.	78,500
Frank P. Palantoni	60,000
Christopher J. Sobecki	40,000
Judith L. Swain, M.D.	40,000
Kathleen M. Wiltsey	40,000

The following table presents the fair value of each grant of stock options in 2008 to non-employee members of our board of directors, computed in accordance with FAS 123(R):

		Number of				
		Securities	curities Exercise Price		Grant Date Fair	
	Grant	Underlying	of Option		Value of	
Name	Date	Options	Awards		Options	
Samuel L. Barker, Ph.D.	4/24/2008	10,000	\$	2.20	\$	15,412
	4/24/2008	10,000	\$	2.20	\$	15,412
Philippe Amouyal	4/24/2008	10,000	\$	2.20	\$	15,412
Raymond Debbane	4/24/2008	10,000	\$	2.20	\$	15,412
Robert J. Lefkowitz, M.D.	4/24/2008	10,000	\$	2.20	\$	15,412
Alan S. Nies, M.D.	4/24/2008	10,000	\$	2.20	\$	15,412
Frank P. Palantoni	4/24/2008	10,000	\$	2.20	\$	15,412
Christopher J. Sobecki	4/24/2008	10,000	\$	2.20	\$	15,412
Judith L. Swain, M.D.	4/24/2008	10,000	\$	2.20	\$	15,412
Kathleen M. Wiltsey	4/24.2008	10,000	\$	2.20	\$	15,412

(4) Consists of amounts paid to Dr. Lefkowitz for his consulting services.

(5) Consists of amounts paid to Dr. Nies for his consulting services as chairman of our medical advisory board.

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Compensation Committee Report

The compensation committee of our board of directors is responsible for evaluating the performance of management, determining the compensation of our executive and other officers and administering our 2000 Equity Incentive Plan, under which stock option grants and other stock awards may be made to our employees.

In performing these functions, the compensation committee has reviewed and discussed with the management of our company the information set forth above under the heading "Executive and Director Compensation — Compensation Discussion and Analysis." Based upon that review and discussion, the compensation committee has recommended to the board of directors that the information set forth above under the heading "Executive and Director Compensation — Compensation — Compensation Discussion and Analysis" be included in this proxy statement and incorporated by reference into our annual report on Form 10-K for the year ended December 31, 2008.

Compensation Committee

Frank P. Palantoni (chair) Philippe J. Amouyal Alan S. Nies, M.D.

The foregoing compensation committee report shall not be deemed "filed" for purposes of Section 18 of the Securities Exchange Act of 1934, or otherwise subject to the liabilities of that section, nor shall it be deemed incorporated by reference by any general statement incorporating by reference this proxy statement into any filing under the Securities Act of 1933 or under the Securities Exchange Act of 1934, except to the extent that we specifically incorporate this compensation committee report by reference.

PROPOSALS OF STOCKHOLDERS

In order for a stockholder proposal to be considered for inclusion in our proxy statement for next year's annual meeting, we must receive the written proposal at our principal executive offices no later than November 13, 2009. Any such proposal must also comply with Securities and Exchange Commission regulations regarding the inclusion of stockholder proposals in company-sponsored proxy materials. Similarly, in order for any stockholder proposal to be otherwise raised during next year's annual meeting, we must receive written notice of the proposal, containing the information required by our bylaws, at our principal executive offices no later than November 13, 2009. You may contact the corporate secretary at our principal executive offices for a copy of the relevant bylaw provisions for making stockholder proposals.

FINANCIAL INFORMATION

Our annual report to stockholders, including financial statements, accompanies this proxy statement but does not constitute a part of the proxy solicitation materials. You may obtain, without charge, a copy of our annual report on Form 10-K, including the financial statements and exhibits thereto, by written request to Corporate Communications, Lexicon Pharmaceuticals, Inc., 8800 Technology Forest Place, The Woodlands, Texas 77381.

By order of the board of directors,

/s/ Jeffrey L. Wade

Jeffrey L. Wade

Secretary

March 13, 2009 The Woodlands, Texas

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Appendix A

LEXICON PHARMACEUTICALS, INC. EQUITY INCENTIVE PLAN

This Plan initially was established as the Lexicon Genetics Incorporated 1995 Stock Option Plan (the "1995 Stock Option Plan"), which was adopted by the Board and approved by the Company's stockholders on September 13, 1995. The 1995 Stock Option Plan was subsequently amended and restated in its entirety and renamed the Lexicon Genetics Incorporated 2000 Equity Incentive Plan (the "2000 Equity Incentive Plan"), which was adopted by the Board on February 3, 2000 and approved by the Company's stockholders on March 15, 2000 and May 19, 2004. The 2000 Equity Incentive Plan is hereby amended and restated in its entirety and renamed the Equity Incentive Plan, effective as of its adoption by the Board subject to approval by the Company's stockholders. The terms of this Plan shall supersede the terms of the 1995 Stock Option Plan and the 2000 Equity Incentive Plan in their entirety; provided, however, that nothing herein shall operate or be construed as modifying the terms of an Incentive Stock Option as being a new grant for purpose of Section 424(h) of the Code.

1.

PURPOSES.

(a) ELIGIBLE STOCK AWARD RECIPIENTS. The persons eligible to receive Stock Awards are the Employees, Directors and Consultants of the Company and its Affiliates.

(b) AVAILABLE STOCK AWARDS. The purpose of the Plan is to provide a means by which eligible recipients of Stock Awards may be given an opportunity to benefit from increases in value of the Common Stock through the granting of the following Stock Awards: (i) Incentive Stock Options, (ii) Nonstatutory Stock Options, (iii) Stock Bonus Awards, (iv) Restricted Stock Awards, (v) Phantom Stock Awards and (vi) Stock Appreciation Rights.

(c) GENERAL PURPOSE. The Company, by means of the Plan, seeks to retain the services of the group of persons eligible to receive Stock Awards, to secure and retain the services of new members of this group and to provide incentives for such persons to exert maximum efforts for the success of the Company and its Affiliates.

2.

DEFINITIONS.

(a) "AFFILIATE" means any parent corporation or subsidiary corporation of the Company, whether now or hereafter existing, as those terms are defined in Sections 424(e) and (f), respectively, of the Code.

(b) "BOARD" means the Board of Directors of the Company.

(c) "CODE" means the Internal Revenue Code of 1986, as amended.

(d) "COMMITTEE" means a committee of one or more members of the Board appointed by the Board in accordance with subsection 3(c).

(e) "COMMON STOCK" means the common stock, par value \$.001 per share, of the Company.

(f) "COMPANY" means Lexicon Pharmaceuticals, Inc. a Delaware corporation.

(g) "CONSULTANT" means any person other than a Director or Employee who is engaged by the Company or an Affiliate to render consulting or advisory services and who is compensated for such services.

(h) "CONTINUOUS SERVICE" means that the Participant's service with the Company or an Affiliate, whether as an Employee, Director or Consultant, is not interrupted or terminated. The Participant's Continuous Service shall not be deemed to have terminated merely because of a change in the capacity in which the Participant renders service to the Company or an Affiliate as an Employee, Consultant or Director or a change in the entity for which the Participant renders such service, provided that there is no interruption or termination of the Participant's Continuous Service. For example, a change in status from an Employee of the Company to a Consultant of an Affiliate or a Director will not constitute an interruption of Continuous Service. The Board or the chief executive officer of the Company, in that party's sole discretion, may determine whether Continuous Service shall be considered interrupted in the case of any leave of absence approved by that party, including sick leave, military leave or any other personal leave.

(i) "COVERED EMPLOYEE" means the chief executive officer and the four (4) other highest compensated officers of the Company for whom total compensation is required to be reported to stockholders under the Exchange Act, as determined for purposes of Section 162(m) of the Code.

(j) "DIRECTOR" means a member of the Board of Directors of the Company.

(k) "DISABILITY" means the permanent and total disability of a person within the meaning of Section 22(e)(3) of the Code.

(1) "EMPLOYEE" means any person (which may include a Director) who is employed by the Company or an Affiliate.

(m) "EXCHANGE ACT" means the Securities Exchange Act of 1934, as amended.

(n) "FAIR MARKET VALUE" means, as of any date, the value of the Common Stock determined as follows:

(i) If the Common Stock is listed on any established stock exchange or traded on the Nasdaq Stock Market, the Fair Market Value of a share of Common Stock shall be the closing sales price for such stock (or the closing bid, if no sales were reported) as quoted on such exchange or market (or the exchange or market with the greatest volume of trading in the Common Stock) on the last market trading day prior to the day of determination, as reported in The Wall Street Journal or such other source as the Board deems reliable.

(ii) In the absence of such markets for the Common Stock, the Fair Market Value shall be determined in good faith by the Board in such manner as it deems appropriate and as is consistent with the requirements of section 409A of the Code.

(o) "INCENTIVE STOCK OPTION" means an option to purchase Common Stock that is intended to qualify as an incentive stock option within the meaning of Section 422 of the Code and the regulations promulgated thereunder.

(p) "NON-EMPLOYEE DIRECTOR" means a Director who either (i) is not a current Employee or Officer of the Company or its parent or a subsidiary, does not receive compensation (directly or indirectly) from the Company or its parent for a subsidiary for services rendered as a consultant or in any capacity other than as a Director (except for an amount as to which disclosure would not be required under Item 404(a) of Regulation S-K promulgated pursuant to the Securities Act ("Regulation S-K")), does not possess an interest in any other transaction as to which disclosure would be required under Item 404(b) of Regulation S-K; or (ii) is otherwise considered a "non-employee director" for purposes of Rule 16b-3.

(q) "NONSTATUTORY STOCK OPTION" means an option to purchase Common Stock other than an Incentive Stock Option.

(r) "OFFICER" means a person who is an officer of the Company within the meaning of Section 16 of the Exchange Act and the rules and regulations promulgated thereunder.

(s) "OPTION" means an Incentive Stock Option or a Nonstatutory Stock Option granted pursuant to Section 6 of the Plan.

(t) "OPTION AGREEMENT" means a written agreement between the Company and an Optionholder evidencing the terms and conditions of an individual Option grant. Each Option Agreement shall be subject to the terms and conditions of the Plan.

(u) "OPTIONHOLDER" means a person to whom an Option is granted pursuant to the Plan or, if applicable, such other person who holds an outstanding Option.

(v) "OUTSIDE DIRECTOR" means a Director who either (i) is not a current employee of the Company or an "affiliated corporation" (within the meaning of Treasury Regulations promulgated under Section 162(m) of the Code), is not a former employee of the Company or an "affiliated corporation" receiving compensation for prior services (other than benefits under a tax qualified pension plan), was not an officer of the Company or an "affiliated corporation" at any time and is not currently receiving direct or indirect remuneration from the Company or an "affiliated corporation" for services in any capacity other than as a Director or (ii) is otherwise considered an "outside director" for purposes of Section 162(m) of the Code.

(w) "PARTICIPANT" means a person to whom a Stock Award is granted pursuant to the Plan or, if applicable, such other person who holds an outstanding Stock Award.

(x) "PHANTOM STOCK AWARD" means a right to receive shares of Common Stock (or the Fair Market Value thereof) granted pursuant to Section 7(c) of the Plan.

(y) "PLAN" means this Lexicon Pharmaceuticals, Inc. Equity Incentive Plan.

(z) "RESTRICTED STOCK AWARD" means a right to purchase restricted Common Stock granted pursuant to Section 7(b) of the Plan.

(aa) "RULE 16B-3" means Rule 16b-3 promulgated under the Exchange Act or any successor to Rule 16b-3, as in effect from time to time.

(bb) "SECURITIES ACT" means the Securities Act of 1933, as amended.

(cc) "STOCK APPRECIATION RIGHT" means a right to receive an amount equal to any appreciation or increase in the Fair Market Value of Common Stock over a specified period of time granted pursuant to Section 7(d) of the Plan.

(dd) "STOCK AWARD" means any right granted under the Plan, including an Option, a Stock Bonus Award, a Restricted Stock Award, a Phantom Stock Award, or a Stock Appreciation Right.

(ee) "STOCK AWARD AGREEMENT" means a written agreement between the Company and a holder of a Stock Award evidencing the terms and conditions of an individual Stock Award grant. Each Stock Award Agreement shall be subject to the terms and conditions of the Plan.

(ff) "STOCK BONUS AWARD" means an award of Common Stock granted pursuant to Section 7(a) of the Plan.

(gg) "TEN PERCENT STOCKHOLDER" means a person who owns (or is deemed to own pursuant to Section 424(d) of the Code) stock possessing more than ten percent (10%) of the total combined voting power of all classes of stock of the Company or of any of its Affiliates.

3.

ADMINISTRATION.

(a) ADMINISTRATION BY BOARD. The Board shall administer the Plan unless and until the Board delegates administration to a Committee, as provided in subsection 3(c).

(b) POWERS OF BOARD. The Board shall have the power, subject to, and within the limitations of, the express provisions of the Plan:

(i) To determine from time to time which of the persons eligible under the Plan shall be granted Stock Awards; when and how each Stock Award shall be granted; what type or combination of types of Stock Award shall be granted; the provisions of each Stock Award granted (which need not be identical), including the time or times when a person shall be permitted to receive Common Stock pursuant to a Stock Award; and the number of shares of Common Stock with respect to which a Stock Award shall be granted to each such person.

(ii) To construe and interpret the Plan and Stock Awards granted under it, and to establish, amend and revoke rules and regulations for its administration. The Board, in the exercise of this power, may correct any defect, omission or inconsistency in the Plan or in any Stock Award Agreement, in a manner and to the extent it shall deem necessary or expedient to make the Plan fully effective.

(iii) To amend the Plan or a Stock Award as provided in Section 12.

(iv) To terminate or suspend the Plan as provided in Section 13.

(v) Generally, to exercise such powers and to perform such acts as the Board deems necessary or expedient to promote the best interests of the Company that are not in conflict with the provisions of the Plan.

(c) DELEGATION TO COMMITTEE.

(i) GENERAL. The Board may delegate administration of the Plan to a Committee or Committees of one (1) or more members of the Board, and the term "Committee" shall apply to any person or persons to whom such authority has been delegated. If administration is delegated to a Committee, the Committee shall have, in connection with the administration of the Plan, the powers theretofore possessed by the Board, including the power to delegate to a subcommittee any of the administrative powers the Committee is authorized to exercise (and references in this Plan to the Board shall thereafter be to the Committee or subcommittee), subject, however, to such resolutions, not inconsistent with the provisions of the Plan, as may be adopted from time to time by the Board. The Board may abolish the Committee at any time and revest in the Board the administration of the Plan.

(ii) COMMITTEE COMPOSITION WHEN COMMON STOCK IS PUBLICLY TRADED. At such time as the Common Stock is publicly traded, in the discretion of the Board, a Committee may consist solely of two or more Outside Directors, in accordance with Section 162(m) of the Code, and/or solely of two or more Non-Employee Directors, in accordance with Rule 16b-3. Within the scope of such authority, the Board or the Committee may (A) delegate to a committee of one or more members of the Board who are not Outside Directors the authority to grant Stock Awards to eligible persons who are either (1) not then Covered Employees and are not expected to be Covered Employees at the time of recognition of income resulting from such Stock Award or (2) not persons with respect to whom the Company wishes to comply with Section 162(m) of the Code, and/or (B) delegate to a committee of one or more members of the Board who are not Stock Awards to eligible persons who are not Non-Employee Directors the authority to grant Stock Awards to eligible persons who are not Non-Employee Directors the authority to grant Stock Awards to eligible persons who are not Non-Employee Directors the authority to grant Stock Awards to eligible persons who are not Non-Employee Directors the authority to grant Stock Awards to eligible persons who are not then subject to Section 16 of the Exchange Act.

(d) EFFECT OF BOARD'S DECISION. All determinations, interpretations and constructions made by the Board in good faith shall not be subject to review by any person and shall be final, binding and conclusive on all persons.

4.

SHARES SUBJECT TO THE PLAN.

(a) SHARE RESERVE. Subject to the provisions of Section 11 relating to adjustments upon changes in Common Stock, the Common Stock that may be issued pursuant to Stock Awards shall not exceed in the aggregate thirty-five million (35,000,000) shares.

(b) REVERSION OF SHARES TO THE SHARE RESERVE. If any Stock Award shall for any reason expire or otherwise terminate, in whole or in part, without having been exercised in full or shares of Common Stock issued to a Participant pursuant to a Stock Award are forfeited to or repurchased by the Company, including any repurchase or forfeiture caused by the failure to meet a contingency or condition required for the vesting of such shares, the shares of Common Stock not issued under such Stock Award or forfeited to or repurchased by the Company shall revert to and again become available for issuance under the Plan; provided, however, that shares subject to a Stock Award that are not delivered to a Participant because (i) such Participant's right to purchase such shares subject to an Option are surrendered in payment of the exercise price for other shares subject to such Option in a "net exercise," or (ii) such shares are withheld in satisfaction of the withholding of taxes incurred in connection with the exercise of an Option or Stock Award, the shares so surrendered or withheld shall not remain available for subsequent issuance under the Plan.

(c) SOURCE OF SHARES. The shares of Common Stock subject to the Plan may be unissued shares or reacquired shares, bought on the market or otherwise.

(d) SHARES AVAILABLE FOR SPECIFIC STOCK AWARDS. Subject to the provisions of Section 11 relating to adjustments upon changes in the shares of Common Stock, the Common Stock that may be issued pursuant to Stock Awards other than Options and Stock Appreciation Rights shall not exceed in the aggregate three million, five hundred thousand (3,500,000) shares.

5.

ELIGIBILITY.

(a) ELIGIBILITY FOR SPECIFIC STOCK AWARDS. Incentive Stock Options may be granted only to Employees. Stock Awards other than Incentive Stock Options may be granted to Employees, Directors and Consultants.

(b) TEN PERCENT STOCKHOLDERS. A Ten Percent Stockholder shall not be granted an Incentive Stock Option unless the exercise price of such Option is at least one hundred ten percent (110%) of the Fair Market Value of the Common Stock at the date of grant and the Option is not exercisable after the expiration of five (5) years from the date of grant.

(c) SECTION 162(m) LIMITATION. Subject to the provisions of Section 11 relating to adjustments upon changes in the shares of Common Stock, no Employee shall be eligible to be granted Options covering more than three million (3,000,000) shares during any calendar year.

6.

OPTION PROVISIONS.

Each Option shall be in such form and shall contain such terms and conditions as the Board shall deem appropriate. All Options shall be separately designated Incentive Stock Options or Nonstatutory Stock Options at the time of grant, and, if certificates are issued, a separate certificate or certificates will be issued for shares of Common Stock

purchased on exercise of each type of Option. The provisions of separate Options need not be identical, but each Option shall include (through incorporation of provisions hereof by reference in the Option or otherwise) the substance of each of the following provisions:

(a) TERM. Subject to the provisions of subsection 5(b) regarding Ten Percent Stockholders, no Option shall be exercisable after the expiration of ten (10) years from the date it was granted.

(b) EXERCISE PRICE. Subject to the provisions of subsection 5(b) regarding Ten Percent Stockholders, the exercise price of each Option shall be not less than one hundred percent (100%) of the Fair Market Value of the Common Stock subject to the Option on the date the Option is granted. Notwithstanding the foregoing, an Option may be granted with an exercise price lower than that set forth in the preceding sentence if such Option is granted pursuant to an assumption or substitution for another option in a manner satisfying the provisions of Section 424(a) of the Code.

(c) CONSIDERATION. The purchase price of Common Stock acquired pursuant to an Option shall be paid, to the extent permitted by applicable statutes and regulations, either (i) in cash at the time the Option is exercised or (ii) at the discretion of the Board (1) by delivery to the Company of other Common Stock, (2) according to a deferred payment or other similar arrangement with the Optionholder, (3) by surrender of Optionholder's right to purchase shares subject to an Option (valued, for such purposes, as the Fair Market Value of such surrendered shares on the date of exercise less the exercise price for such surrendered shares) in payment of the exercise price for other shares subject to such Option in a "net exercise" of such Option, or (4) in any other form of legal consideration that may be acceptable to the Board. At any time that the Company is incorporated in Delaware, payment of the Common Stock's "par value," as defined in the Delaware General Corporation Law, shall not be made by deferred payment. In the case of any deferred payment arrangement, interest shall be compounded at least annually and shall be charged at the minimum rate of interest necessary to avoid the treatment as interest, under any applicable provisions of the Code, of any amounts other than amounts stated to be interest under the deferred payment arrangement.

(d) TRANSFERABILITY. An Incentive Stock Option shall not be transferable except by will or by the laws of descent and distribution and shall be exercisable during the lifetime of the Optionholder only by the Optionholder. A Nonstatutory Stock Option shall be transferable to the extent provided in the Option Agreement; provided that, if the Nonstatutory Stock Option does not provide for transferability, then the Nonstatutory Stock Option shall not be transferable except by will or by the laws of descent and distribution and shall be exercisable during the lifetime of the Optionholder only by the Optionholder. Notwithstanding the foregoing, the Optionholder may, by delivering written notice to the Company, in a form satisfactory to the Company, designate a third party who, in the event of the death of the Optionholder, shall thereafter be entitled to exercise the Option.

(e) VESTING GENERALLY. The total number of shares of Common Stock subject to an Option may, but need not, vest and therefore become exercisable in periodic installments that may, but need not, be equal. The Option may be subject to such other terms and conditions on the time or times when it may be exercised (which may be based on performance or other criteria) as the Board may deem appropriate. The vesting provisions of individual Options may vary.

(f) TERMINATION OF CONTINUOUS SERVICE. In the event an Optionholder's Continuous Service terminates (other than upon the Optionholder's death or Disability), the Optionholder may exercise his or her Option (to the extent that the Optionholder was entitled to exercise such Option as of the date of termination) but only within such period of time ending on the earlier of (i) the date three (3) months following the termination of the Optionholder's Continuous Service (or such longer or shorter period specified in the Option Agreement), or (ii) the expiration of the term of the Option as set forth in the Option Agreement. If, after termination, the Optionholder does not exercise his or her Option within the time specified in the Option Agreement, the Option shall terminate.

(g) EXTENSION OF TERMINATION DATE. An Optionholder's Option Agreement may also provide that if the exercise of the Option following the termination of the Optionholder's Continuous Service (other than upon the Optionholder's death or Disability) would be prohibited at any time solely because the issuance of shares of Common Stock would violate the registration requirements under the Securities Act, then the Option shall terminate on the earlier of (i) the expiration of the term of the Option set forth in subsection 6(a) or (ii) the expiration of a period of three (3) months after the termination of the Optionholder's Continuous Service during which the exercise of the Option would not be in violation of such registration requirements.

(h) DISABILITY OF OPTIONHOLDER. In the event that an Optionholder's Continuous Service terminates as a result of the Optionholder's Disability, the Optionholder may exercise his or her Option (to the extent that the Optionholder was entitled to exercise such Option as of the date of termination), but only within such period of time ending on the earlier of (i) the date twelve (12) months following such termination (or such longer or shorter period specified in the Option Agreement,) or (ii) the expiration of the term of the Option as set forth in the Option Agreement. If, after termination, the Optionholder does not exercise his or her Option within the time specified herein, the Option shall terminate.

(i) DEATH OF OPTIONHOLDER. In the event (i) an Optionholder's Continuous Service terminates as a result of the Optionholder's death or (ii) the Optionholder dies within the period (if any) specified in the Option Agreement after the termination of the Optionholder's Continuous Service for a reason other than death, then the Option may be exercised (to the extent the Optionholder was entitled to exercise such Option as of the date of death) by the Optionholder's estate, by a person who acquired the right to exercise the Option by bequest or inheritance or by a person designated to exercise the Option upon the Optionholder's death pursuant to subsection 6(d), but only within the period ending on the earlier of (1) the date eighteen (18) months following the date of death (or such longer or shorter period specified in the Option Agreement) or (2) the expiration of the term of such Option as set forth in the Option Agreement. If, after death, the Option is not exercised within the time specified herein, the Option shall terminate.

PROVISIONS OF STOCK AWARDS OTHER THAN OPTIONS.

7.

(a) STOCK BONUS AWARDS. Each Stock Bonus Award agreement shall be in such form and shall contain such terms and conditions as the Board shall deem appropriate. The terms and conditions of Stock Bonus Award agreements may change from time to time, and the terms and conditions of separate Stock Bonus Award agreements need not be identical, but each Stock Bonus Award agreement shall include (through incorporation of provisions hereof by reference in the agreement or otherwise) the substance of each of the following provisions:

(i) CONSIDERATION. A Stock Bonus Award may be granted in consideration for past services actually rendered to the Company or an Affiliate for its benefit.

(ii) VESTING. Shares of Common Stock awarded under the Stock Bonus Award agreement may, but need not, be subject to a share repurchase option or forfeiture restrictions in favor of the Company in accordance with a vesting schedule to be determined by the Board.

(iii) TERMINATION OF PARTICIPANT'S CONTINUOUS SERVICE. In the event a Participant's Continuous Service terminates, the Company may reacquire any or all of the shares of Common Stock held by the Participant which have not vested as of the date of termination under the terms of the Stock Bonus Award agreement.

(iv) TRANSFERABILITY. Rights to acquire shares of Common Stock under the Stock Bonus Award agreement shall be transferable by the Participant only upon such terms and conditions as are set forth in the Stock Bonus Award agreement, as the Board shall determine in its discretion, so long as Common Stock awarded under the Stock Bonus Award agreement remains subject to the terms of the Stock Bonus Award agreement.

(b) RESTRICTED STOCK AWARDS. Each Restricted Stock Award agreement shall be in such form and shall contain such terms and conditions as the Board shall deem appropriate. The terms and conditions of the Restricted Stock Award agreement may change from time to time, and the terms and conditions of separate Restricted Stock Award agreements need not be identical, but each Restricted Stock Award agreement shall include (through incorporation of provisions hereof by reference in the agreement or otherwise) the substance of each of the following provisions:

(i) PURCHASE PRICE. The purchase price under each Restricted Stock Award agreement shall be such amount as the Board shall determine and designate in such Restricted Stock Award agreement. Such purchase price shall not be less than eighty-five percent (85%) of the Common Stock's Fair Market Value on the date such award is made or at the time the purchase is consummated.

(ii) CONSIDERATION. The purchase price of Common Stock acquired pursuant to the Restricted Stock Award agreement shall be paid either: (A) in cash at the time of purchase; (B) at the discretion of the Board, according to a deferred payment or other similar arrangement with the Participant; or (C) in any other form of legal consideration that may be acceptable to the Board in its discretion; provided, however, that at any time that the Company is incorporated in Delaware, then payment of the Common Stock's "par value," as defined in the Delaware General Corporation Law, shall not be made by deferred payment.

(iii) VESTING. Shares of Common Stock acquired under the Restricted Stock Award agreement may, but need not, be subject to a share repurchase option in favor of the Company in accordance with a vesting schedule to be determined by the Board.

(iv) TERMINATION OF PARTICIPANT'S CONTINUOUS SERVICE. In the event a Participant's Continuous Service terminates, the Company may repurchase or otherwise reacquire any or all of the shares of Common Stock held by the Participant which have not vested as of the date of termination under the terms of the Restricted Stock Award agreement.

(v) TRANSFERABILITY. Rights to acquire shares of Common Stock under the Restricted Stock Award agreement shall be transferable by the Participant only upon such terms and conditions as are set forth in the Restricted Stock Award agreement, as the Board shall determine in its discretion, so long as Common Stock awarded under the Restricted Stock Award agreement remains subject to the terms of the Restricted Stock Award agreement.

(c) PHANTOM STOCK AWARDS. Each Phantom Stock Award agreement shall be in such form and shall contain such terms and conditions as the Board shall deem appropriate. The terms and conditions of Phantom Stock Award agreements may change from time to time, and the terms and conditions of separate Phantom Stock Award agreements need not be identical, provided, however, that each Phantom Stock Award agreement shall include (through incorporation of the provisions hereof by reference in the agreement or otherwise) the substance of each of the following provisions:

(i) CONSIDERATION. At the time of grant of a Phantom Stock Award, the Board will determine the consideration, if any, to be paid by the Participant upon delivery of each share of Common Stock subject to the Phantom Stock Award. The consideration to be paid (if any) by the Participant for each share of Common Stock subject to a Phantom Stock Award may be paid in any form of legal consideration that may be acceptable to the Board in its sole discretion and permissible under applicable law.

(ii) VESTING. At the time of the grant of a Phantom Stock Award, the Board may impose such restrictions or conditions to the vesting of the Phantom Stock Award as it, in its sole discretion, deems appropriate.

(iii) PAYMENT. A Phantom Stock Award may be settled by the delivery of shares of Common Stock, their cash equivalent, any combination thereof or in any other form of consideration, as determined by the Board and contained in the Phantom Stock Award agreement.

(iv) DIVIDEND EQUIVALENTS. Dividend equivalents may be credited in respect of shares of Common Stock covered by a Phantom Stock Award, as determined by the Board and contained in the Phantom Stock Award agreement. At the sole discretion of the Board, such dividend equivalents may be converted into additional shares of Common Stock covered by the Phantom Stock Award in such manner as determined by the Board. Any additional shares covered by the Phantom Stock Award credited by reason of such dividend equivalents will be subject to all the terms and conditions of the underlying Phantom Stock Award agreement to which they relate.

(v) TERMINATION OF PARTICIPANT'S CONTINUOUS SERVICE. Except as otherwise provided in the applicable Phantom Stock Award agreement, such portion of the Phantom Stock Award that has not vested will be forfeited upon the Participant's termination of Continuous Service.

(vi) TRANSFERABILITY. Rights under the Phantom Stock Award agreement shall be transferable by the Participant only upon such terms and conditions as are set forth in the Phantom Stock Award agreement, as the Board shall determine in its discretion.

(d) STOCK APPRECIATION RIGHTS. Each Stock Appreciation Right agreement shall be in such form and shall contain such terms and conditions as the Board shall deem appropriate. The terms and conditions of the Stock Appreciation Right agreements may change from time to time, and the terms and conditions of separate Stock Appreciation Right agreements need not be identical, but each Stock Appreciation Right agreement shall include (through incorporation of provisions hereof by reference in the agreement or otherwise) the substance of each of the following provisions:

(i) CALCULATION OF APPRECIATION. Each Stock Appreciation Right will be denominated in shares of Common Stock equivalents. The appreciation distribution payable on the exercise of a Stock Appreciation Right will be not greater than an amount equal to the excess of (i) the aggregate Fair Market Value (on the date of the exercise of the Stock Appreciation Right) of a number of shares of Common Stock equal to the number of shares of Common Stock equivalents in which the Participant is vested under such Stock Appreciation Right, and with respect to which the Participant is exercising the Stock Appreciation Right on such date, over (ii) an amount (the strike price) that will be determined by the Board at the time of grant of the Stock Appreciation Right for such number of shares of Common Stock, provided that the strike price of a Stock Appreciation Right shall be not less than one hundred percent (100%) of the Fair Market Value of the Common Stock equal to the number of shares of Common Stock equivalents subject to the Stock Appreciation Right on the date the Stock Appreciation Right is granted.

(ii) VESTING. At the time of the grant of a Stock Appreciation Right, the Board may impose such restrictions or conditions to the vesting of such Stock Appreciation Right as it, in its sole discretion, deems appropriate.

(iii) EXERCISE. To exercise any outstanding Stock Appreciation Right, the Participant must provide written notice of exercise to the Company in compliance with the provisions of the Stock Appreciation Right agreement evidencing such Stock Appreciation Right.

(iv) PAYMENT. The appreciation distribution in respect to a Stock Appreciation Right may be paid in cash, shares of Common Stock, a combination of cash and shares of Common Stock or in any other form of consideration, as determined by the Board and contained in the Stock Appreciation Right agreement evidencing such Stock Appreciation Right.

(v) TERMINATION OF PARTICIPANT'S CONTINUOUS SERVICE. In the event that a Participant's Continuous Service terminates, the Participant may exercise his or her Stock Appreciation Right (to the extent that the Participant was entitled to exercise such Stock Appreciation Right as of the date of termination) but only within such period of time ending on the earlier of (i) the date three (3) months following the termination of the Participant's Continuous Service (or such longer or shorter period specified in the Stock Appreciation Right agreement), or (ii) the expiration of the term of the Stock Appreciation Right as set forth in the Stock Appreciation Right agreement. If, after termination, the Participant does not exercise his or her Stock Appreciation Right within the time specified herein or in the Stock Appreciation Right agreement (as applicable), the Stock Appreciation Right shall terminate.

(vi) TRANSFERABILITY. Rights under the Stock Appreciation Right agreement shall be transferable by the Participant only upon such terms and conditions as are set forth in the Stock Appreciation Right agreement, as the Board shall determine in its discretion.

8.

COVENANTS OF THE COMPANY.

(a) AVAILABILITY OF SHARES. During the terms of the Stock Awards, the Company shall keep available at all times the number of shares of Common Stock required to satisfy such Stock Awards.

(b) SECURITIES LAW COMPLIANCE. The Company shall seek to obtain from each regulatory commission or agency having jurisdiction over the Plan such authority as may be required to grant Stock Awards and to issue and sell shares of Common Stock upon exercise of the Stock Awards; provided, however, that this undertaking shall not require the Company to register under the Securities Act the Plan, any Stock Award or any Common Stock issued or issuable pursuant to any such Stock Award. If, after reasonable efforts, the Company is unable to obtain from any such regulatory commission or agency the authority which counsel for the Company deems necessary for the lawful issuance and sale of Common Stock under the Plan, the Company shall be relieved from any liability for failure to issue and sell Common Stock upon exercise of such Stock Awards unless and until such authority is obtained.

9.

USE OF PROCEEDS FROM STOCK.

Proceeds from the sale of Common Stock pursuant to Stock Awards shall constitute general funds of the Company.

10.

MISCELLANEOUS.

(a) ACCELERATION OF EXERCISABILITY AND VESTING. The Board shall have the power to accelerate the time at which a Stock Award may first be exercised or the time during which a Stock Award or any part thereof will vest in accordance with the Plan, notwithstanding the provisions in the Stock Award stating the time at which it may first be exercised or the time during which it will vest.

(b) STOCKHOLDER RIGHTS. No Participant shall be deemed to be the holder of, or to have any of the rights of a holder with respect to, any shares of Common Stock subject to such Stock Award unless and until such Participant has satisfied all requirements for exercise of the Stock Award pursuant to its terms.

(c) NO EMPLOYMENT OR OTHER SERVICE RIGHTS. Nothing in the Plan or any instrument executed or Stock Award granted pursuant thereto shall confer upon any Participant any right to continue to serve the Company or an Affiliate in the capacity in effect at the time the Stock Award was granted or shall affect the right of the Company or an Affiliate to terminate (i) the employment of an Employee with or without notice and with or without cause, (ii) the service of a Consultant pursuant to the terms of such Consultant's agreement with the Company or an Affiliate or (iii) the service of a Director pursuant to the Bylaws of the Company or an Affiliate, and any applicable provisions of the corporate law of the state in which the Company or the Affiliate is incorporated, as the case may be. A-10

(d) INCENTIVE STOCK OPTION \$100,000 LIMITATION. To the extent that the aggregate Fair Market Value (determined at the time of grant) of Common Stock with respect to which Incentive Stock Options are exercisable for the first time by any Optionholder during any calendar year (under all plans of the Company and its Affiliates) exceeds one hundred thousand dollars (\$100,000), the Options or portions thereof which exceed such limit (according to the order in which they were granted) shall be treated as Nonstatutory Stock Options.

(e) INVESTMENT ASSURANCES. The Company may require a Participant, as a condition of exercising or acquiring Common Stock under any Stock Award, (i) to give written assurances satisfactory to the Company as to the Participant's knowledge and experience in financial and business matters and/or to employ a purchaser representative reasonably satisfactory to the Company who is knowledgeable and experienced in financial and business matters and that he or she is capable of evaluating, alone or together with the purchaser representative, the merits and risks of exercising the Stock Award; and (ii) to give written assurances satisfactory to the Company stating that the Participant is acquiring Common Stock subject to the Stock Award for the Participant's own account and not with any present intention of selling or otherwise distributing the Common Stock. The foregoing requirements, and any assurances given pursuant to such requirements, shall be inoperative if (1) the issuance of the shares of Common Stock upon the exercise or acquisition of Common Stock under the Stock Award has been registered under a then currently effective registration statement under the Securities Act or (2) as to any particular requirement, a determination is made by counsel for the Company that such requirement need not be met in the circumstances under the then applicable securities laws. The Company may, upon advice of counsel to the Company, place legends on stock certificates issued under the Plan as such counsel deems necessary or appropriate in order to comply with applicable securities laws, including, but not limited to, legends restricting the transfer of the Common Stock.

(f) WITHHOLDING OBLIGATIONS. To the extent provided by the terms of a Stock Award Agreement, the Participant may satisfy any federal, state or local tax withholding obligation relating to the exercise or acquisition of Common Stock under a Stock Award by any of the following means (in addition to the Company's right to withhold from any compensation paid to the Participant by the Company) or by a combination of such means: (i) tendering a cash payment; (ii) authorizing the Company to withhold shares of Common Stock from the shares of Common Stock otherwise issuable to the Participant as a result of the exercise or acquisition of Common Stock under the Stock Award, provided, however, that no shares of Common Stock are withheld with a value exceeding the minimum amount of tax required to be withheld by law; or (iii) delivering to the Company owned and unencumbered shares of Common Stock.

11.

ADJUSTMENTS UPON CHANGES IN STOCK.

(a) CAPITALIZATION ADJUSTMENTS. If any change is made in the Common Stock subject to the Plan, or subject to any Stock Award, without the receipt of consideration by the Company (through merger, consolidation, reorganization, recapitalization, reincorporation, stock dividend, dividend in property other than cash, stock split, liquidating dividend, combination of shares, exchange of shares, change in corporate structure or other transaction not involving the receipt of consideration by the Company), the Plan will be appropriately adjusted in the class(es) and the number of securities subject to the Plan pursuant to subsection 4(a), the maximum number of securities subject to Stock Awards other than Options and Stock Appreciation Rights pursuant to subsection 4(d), the maximum number of securities subject to award to any person pursuant to subsection 5(c), and the outstanding Stock Awards will be appropriately adjusted in the class(es) and number of securities and price per share of Common Stock subject to such outstanding Stock Awards. The Board shall make such adjustments, and its determination shall be final, binding and conclusive. For clarity, the conversion of any convertible securities of the Company shall not be treated as a transaction "without receipt of consideration" by the Company.

(b) DISSOLUTION OR LIQUIDATION. In the event of a dissolution or liquidation of the Company, then all outstanding Stock Awards shall terminate immediately prior to such event.

(c) ASSET SALE, MERGER, CONSOLIDATION OR REVERSE MERGER. In the event of (i) a sale, lease or other disposition of all or substantially all of the assets of the Company, (ii) a merger or consolidation in which the Company is not the surviving corporation or (iii) a reverse merger in which the Company is the surviving corporation but the shares of Common Stock outstanding immediately preceding the merger are converted by virtue of the merger into other property, whether in the form of securities, cash or otherwise, then any surviving corporation or acquiring corporation shall assume any Stock Awards outstanding under the Plan or shall substitute similar stock awards (including an award to acquire the same consideration paid to the stockholders in the transaction described in this subsection 11(c) for those outstanding under the Plan). In the event any surviving corporation or acquiring corporation fails to assume such Stock Awards or to substitute similar stock awards for those outstanding under the Plan, then with respect to Stock Awards held by Participants whose Continuous Service has not terminated, the vesting of such Stock Awards (and, if applicable, the time during which such Stock Awards may be exercised) shall be accelerated in full, and the Stock Awards shall terminate if not exercised (if applicable) at or prior to such event. With respect to any other Stock Awards outstanding under the Plan, such Stock Awards shall terminate if not exercised (if applicable) at or prior to such event.

12. AMENDMENT OF THE PLAN AND STOCK AWARDS.

(a) AMENDMENT OF PLAN. The Board at any time, and from time to time, may amend the Plan. However, except as provided in Section 11 relating to adjustments upon changes in Common Stock, no amendment shall be effective unless approved by the stockholders of the Company to the extent stockholder approval is necessary to satisfy the requirements of Section 422 of the Code, Rule 16b-3 or any Nasdaq or securities exchange listing requirements.

(b) STOCKHOLDER APPROVAL. The Board may, in its sole discretion, submit any other amendment to the Plan for stockholder approval, including, but not limited to, amendments to the Plan intended to satisfy the requirements of Section 162(m) of the Code and the regulations thereunder regarding the exclusion of performance-based compensation from the limit on corporate deductibility of compensation paid to certain executive officers.

(c) CONTEMPLATED AMENDMENTS. It is expressly contemplated that the Board may amend the Plan in any respect the Board deems necessary or advisable to provide eligible Employees with the maximum benefits provided or to be provided under the provisions of the Code and the regulations promulgated thereunder relating to Incentive Stock Options and/or to bring the Plan and/or Incentive Stock Options granted under it into compliance therewith.

(d) NO IMPAIRMENT OF RIGHTS. Rights under any Stock Award granted before amendment of the Plan shall not be impaired by any amendment of the Plan unless (i) the Company requests the consent of the Participant and (ii) the Participant consents in writing.

(e) AMENDMENT OF STOCK AWARDS. The Board at any time, and from time to time, may amend the terms of any one or more Stock Awards; provided, however, that the rights under any Stock Award shall not be impaired by any such amendment unless (i) the Company requests the consent of the Participant and (ii) the Participant consents in writing.

13. TERMINATION OR SUSPENSION OF THE PLAN.

(a) PLAN TERM. The Board may suspend or terminate the Plan at any time. Unless sooner terminated, the Plan shall terminate on the day before the tenth (10th) anniversary of the date the Plan is adopted by the Board or approved

by the stockholders of the Company, whichever is earlier. No Stock Awards may be granted under the Plan while the Plan is suspended or after it is terminated.

(b) NO IMPAIRMENT OF RIGHTS. Suspension or termination of the Plan shall not impair rights and obligations under any Stock Award granted while the Plan is in effect except with the written consent of the Participant.

14. EFFECTIVE DATE OF PLAN.

The Plan shall become effective upon its adoption by the Board, but no Stock Award shall be exercised (or, in the case of a stock bonus, shall be granted) unless and until the Plan has been approved by the stockholders of the Company, which approval shall be within twelve (12) months before or after the date the Plan is adopted by the Board.

15.

CHOICE OF LAW.

The law of the State of Delaware shall govern all questions concerning the construction, validity and interpretation of this Plan, without regard to such state's conflict of laws rules.

Appendix B

LEXICON PHARMACEUTICALS, INC. NON-EMPLOYEE DIRECTORS' STOCK OPTION PLAN

This Plan initially was established as the 2000 Non-Employee Directors' Stock Option Plan, effective as of April 12, 2000 (the "Initial Plan"). The Initial Plan is hereby amended and restated in its entirety and renamed the Non-Employee Directors' Stock Option Plan, effective as of its adoption by the Board subject to approval by the Company's stockholders. The terms of this Plan shall supersede the terms of Initial Plan in their entirety.

1. PURPOSES.

(a) ELIGIBLE OPTION RECIPIENTS. The persons eligible to receive Options are the Non-Employee Directors of the Company.

(c) AVAILABLE OPTIONS. The purpose of the Plan is to provide a means by which Non-Employee Directors may be given an opportunity to benefit from increases in value of the Common Stock through the granting of Nonstatutory Stock Options.

(d) GENERAL PURPOSE. The Company, by means of the Plan, seeks to retain the services of its Non-Employee Directors, to secure and retain the services of new Non-Employee Directors and to provide incentives for such persons to exert maximum efforts for the success of the Company and its Affiliates.

2. DEFINITIONS.

(a) "AFFILIATE" means any parent corporation or subsidiary corporation of the Company, whether now or hereafter existing, as those terms are defined in Sections 424(e) and (f), respectively, of the Code.

(b) "ANNUAL GRANT" means an Option granted annually to all Non-Employee Directors who meet the specified criteria pursuant to subsection 6(b) of the Plan.

(c) "ANNUAL MEETING" means the annual meeting of the stockholders of the Company.

(d) "BOARD" means the Board of Directors of the Company.

(e) "CODE" means the Internal Revenue Code of 1986, as amended.

(f) "COMMON STOCK" means the common stock, par value \$.001 per share, of the Company.

(g) "COMPANY" means Lexicon Pharmaceuticals, Inc., a Delaware corporation.

(h) "CONSULTANT" means any person other than a Director or Employee who is engaged by the Company or an Affiliate to render consulting or advisory services and who is compensated for such services.

(i) "CONTINUOUS SERVICE" means that the Optionholder's service with the Company or an Affiliate, whether as an Employee, Director or Consultant, is not interrupted or terminated. The Optionholder's Continuous Service shall not be deemed to have terminated merely because of a change in the capacity in which the Optionholder renders service to the Company or an Affiliate as an Employee, Consultant or Director or a change in the entity for which the Optionholder renders such service, provided that there is no interruption or termination of the Optionholder's Continuous Service. For example, a change in status from a Non-Employee Director of the Company to a Consultant

of an Affiliate or an Employee of the Company will not constitute an interruption of Continuous Service. The Board or the chief executive officer of the Company, in that party's sole discretion, may determine whether Continuous Service shall be considered interrupted in the case of any leave of absence approved by that party, including sick leave, military leave or any other personal leave.

(j) "DIRECTOR" means a member of the Board of Directors of the Company.

(k) "DISABILITY" means the permanent and total disability of a person within the meaning of Section 22(e)(3) of the Code.

(1) "EMPLOYEE" means any person employed by the Company or an Affiliate. Mere service as a Director or payment of a director's fee by the Company or an Affiliate shall not be sufficient to constitute "employment" by the Company or an Affiliate.

(m) "EXCHANGE ACT" means the Securities Exchange Act of 1934, as amended.

(n) "FAIR MARKET VALUE" means, as of any date, the value of the Common Stock determined as follows:

(i) If the Common Stock is listed on any established stock exchange or traded on the Nasdaq Stock Market, the Fair Market Value of a share of Common Stock shall be the closing sales price for such stock (or the closing bid, if no sales were reported) as quoted on such exchange or market (or the exchange or market with the greatest volume of trading in the Common Stock) on the last market trading day prior to the day of determination, as reported in The Wall Street Journal or such other source as the Board deems reliable.

(ii) In the absence of such markets for the Common Stock, the Fair Market Value shall be determined in good faith by the Board in such manner as it deems appropriate and as is consistent with the requirements of section 409A of the Code.

(o) "INITIAL GRANT" means an Option granted to a Non-Employee Director who meets the specified criteria pursuant to subsection 6(a) of the Plan.

(p) "NON-EMPLOYEE CHAIRMAN" means a Non-Employee Director serving as chairman of the Board.

(q) "NON-EMPLOYEE DIRECTOR" means a Director who is not an Employee.

(r) "OPTION" means an option to purchase Common Stock granted pursuant to the Plan.

(s) "OPTION AGREEMENT" means a written agreement between the Company and an Optionholder evidencing the terms and conditions of an individual Option grant. Each Option Agreement shall be subject to the terms and conditions of the Plan.

(t) "OPTIONHOLDER" means a person to whom an Option is granted pursuant to the Plan or, if applicable, such other person who holds an outstanding Option.

(u) "PLAN" means this Lexicon Pharmaceuticals, Inc. Non-Employee Directors' Stock Option Plan.

(v) "RULE 16B-3" means Rule 16b-3 promulgated under the Exchange Act or any successor to Rule 16b-3, as in effect from time to time.

(w) "SECURITIES ACT" means the Securities Act of 1933, as amended.

3. ADMINISTRATION.

(a) ADMINISTRATION BY BOARD. The Board shall administer the Plan. The Board may not delegate administration of the Plan to a committee.

(b) POWERS OF BOARD. The Board shall have the power, subject to, and within the limitations of, the express provisions of the Plan:

(i) To determine the provisions of each Option to the extent not specified in the Plan.

(ii) To construe and interpret the Plan and Options granted under it, and to establish, amend and revoke rules and regulations for its administration. The Board, in the exercise of this power, may correct any defect, omission or inconsistency in the Plan or in any Option Agreement, in a manner and to the extent it shall deem necessary or expedient to make the Plan fully effective.

(iii) To amend the Plan or an Option as provided in Section 12.

(iv) Generally, to exercise such powers and to perform such acts as the Board deems necessary or expedient to promote the best interests of the Company that are not in conflict with the provisions of the Plan.

(c) EFFECT OF BOARD'S DECISION. All determinations, interpretations and constructions made by the Board in good faith shall not be subject to review by any person and shall be final, binding and conclusive on all persons.

4. SHARES SUBJECT TO THE PLAN.

(a) SHARE RESERVE. Subject to the provisions of Section 11 relating to adjustments upon changes in the Common Stock, the Common Stock that may be issued pursuant to Options shall not exceed in the aggregate one million, two hundred thousand (1,200,000) shares of Common Stock.

(b) REVERSION OF SHARES TO THE SHARE RESERVE. If any Option shall for any reason expire or otherwise terminate, in whole or in part, without having been exercised in full, the shares of Common Stock not acquired under such Option shall revert to and again become available for issuance under the Plan. For clarity, shares subject to an Option that are not delivered to an Optionholder because (i) such Optionholder's right to purchase such shares are surrendered in payment of the exercise price for other shares subject to such Option in a "net exercise," or (ii) such shares are withheld in satisfaction of the withholding of taxes incurred in connection with the exercise of such Option, the shares so surrendered or withheld shall not remain available for subsequent issuance under the Plan.

(c) SOURCE OF SHARES. The shares of Common Stock subject to the Plan may be unissued shares or reacquired shares, bought on the market or otherwise.

5. ELIGIBILITY.

The Options as set forth in section 6 automatically shall be granted under the Plan to all Non-Employee Directors.

6. NON-DISCRETIONARY GRANTS.

(a) INITIAL GRANTS. Without any further action of the Board, each person who is elected or appointed for the first time to be a Non-Employee Director automatically shall, upon the date of his or her initial election or appointment to be a Non-Employee Director, be granted an Initial Grant to purchase Thirty Thousand (30,000) shares of Common Stock on the terms and conditions set forth herein.

(b) ANNUAL GRANTS. Without any further action of the Board, on the day following each Annual Meeting, (i) each person who is then a Non-Employee Director, but is not serving as Non-Employee Chairman, and has been a Non-Employee Director for at least six (6) months, automatically shall be granted an Annual Grant to purchase Ten Thousand (10,000) shares of Common Stock and (ii) the person, if any, who is then Non-Employee Chairman, and has been a Non-Employee Director for at least six (6) months, automatically shall be granted an Annual Option to purchase Twenty Thousand (20,000) shares of Common Stock, each such Option to be granted on the terms and conditions set forth herein.

7. OPTION PROVISIONS.

Each Option shall be in such form and shall contain such terms and conditions as required by the Plan. Each Option shall contain such additional terms and conditions, not inconsistent with the Plan, as the Board shall deem appropriate. Each Option shall include (through incorporation of provisions hereof by reference in the Option or otherwise) the substance of each of the following provisions:

(a) TERM. No Option shall be exercisable after the expiration of ten (10) years from the date it was granted.

(b) EXERCISE PRICE. The exercise price of each Option shall be one hundred percent (100%) of the Fair Market Value of the stock subject to the Option on the date the Option is granted. Notwithstanding the foregoing, an Option may be granted with an exercise price lower than that set forth in the preceding sentence if such Option is granted pursuant to an assumption or substitution for another option in a manner satisfying the provisions of Section 424(a) of the Code.

(c) CONSIDERATION. The purchase price of stock acquired pursuant to an Option may be paid, to the extent permitted by applicable statutes and regulations, in any combination of (i) cash or check, (ii) delivery to the Company of other Common Stock, or (iii) surrender of Optionholder's right to purchase shares subject to such Option (valued, for such purposes, as the Fair Market Value of such surrendered shares on the date of exercise less the exercise price for such surrendered shares) in payment of the exercise price for other shares subject to such Option in a "net exercise" of such Option.

(d) TRANSFERABILITY. An Option is not transferable, except (i) by will or by the laws of descent and distribution, (ii) by instrument to an inter vivos or testamentary trust, in a form accepted by the Company, in which the Option is to be passed to beneficiaries upon the death of the trustor (settlor) and (iii) by gift, in a form accepted by the Company, to a member of the "immediate family" of the Optionholder as that term is defined in 17 C.F.R. 240.16a-1(e). In addition, the Optionholder may, by delivering written notice to the Company, in a form satisfactory to the Company, designate a third party who, in the event of the death of the Optionholder, shall thereafter be entitled to exercise the Option.

(e) VESTING. Options shall vest as follows:

(i) Initial Grants shall provide for vesting of 1/60th of the shares subject to the Option each month after grant for five (5) years after the date of the grant.

(ii) Annual Grants shall provide for vesting of 1/12th of the shares subject to the Option each month after grant for twelve (12) months after the date of the grant.

(f) TERMINATION OF CONTINUOUS SERVICE. In the event an Optionholder's Continuous Service terminates (other than upon the Optionholder's death or Disability), the Optionholder may exercise his or her Option (to the extent that the Optionholder was entitled to exercise it as of the date of termination) but only within such period of time ending on the earlier of (i) the date six (6) months following the termination of the Optionholder's Continuous Service, or (ii) the expiration of the term of the Option as set forth in the Option Agreement. If, after termination, the Optionholder does not exercise his or her Option within the time specified in the Option Agreement, the Option shall terminate.

(g) EXTENSION OF TERMINATION DATE. If the exercise of the Option following the termination of the Optionholder's Continuous Service (other than upon the Optionholder's death or Disability) would be prohibited at any time solely because the issuance of shares would violate the registration requirements under the Securities Act, then the Option shall terminate on the earlier of (i) the expiration of the term of the Option set forth in subsection 7(a) or (ii) the expiration of a period of three (3) months after the termination of the Optionholder's Continuous Service during

which the exercise of the Option would not be in violation of such registration requirements.

(h) DISABILITY OF OPTIONHOLDER. In the event an Optionholder's Continuous Service terminates as a result of the Optionholder's Disability, the Optionholder may exercise his or her Option (to the extent that the Optionholder was entitled to exercise it as of the date of termination), but only within such period of time ending on the earlier of (i) the date twelve (12) months following such termination or (ii) the expiration of the term of the Option as set forth in the Option Agreement. If, after termination, the Optionholder does not exercise his or her Option within the time specified herein, the Option shall terminate.

(i) DEATH OF OPTIONHOLDER. In the event (i) an Optionholder's Continuous Service terminates as a result of the Optionholder's death or (ii) the Optionholder dies within the three-month period after the termination of the Optionholder's Continuous Service for a reason other than death, then the Option may be exercised (to the extent the Optionholder was entitled to exercise the Option as of the date of death) by the Optionholder's estate, by a person who acquired the right to exercise the Option by bequest or inheritance or by a person designated to exercise the Option upon the Optionholder's death, but only within the period ending on the earlier of (A) the date eighteen (18) months following the date of death or (B) the expiration of the term of such Option as set forth in the Option Agreement. If, after death, the Option is not exercised within the time specified herein, the Option shall terminate.

8. COVENANTS OF THE COMPANY.

(a) AVAILABILITY OF SHARES. During the terms of the Options, the Company shall keep available at all times the number of shares of Common Stock required to satisfy such Options.

(b) SECURITIES LAW COMPLIANCE. The Company shall seek to obtain from each regulatory commission or agency having jurisdiction over the Plan such authority as may be required to grant Options and to issue and sell shares of Common Stock upon exercise of the Options; provided, however, that this undertaking shall not require the Company to register under the Securities Act the Plan, any Option or any stock issued or issuable pursuant to any such Option. If, after reasonable efforts, the Company is unable to obtain from any such regulatory commission or agency the authority which counsel for the Company deems necessary for the lawful issuance and sale of stock under the Plan, the Company shall be relieved from any liability for failure to issue and sell stock upon exercise of such Options unless and until such authority is obtained.

9. USE OF PROCEEDS FROM STOCK.

Proceeds from the sale of stock pursuant to Options shall constitute general funds of the Company.

10. MISCELLANEOUS.

(a) STOCKHOLDER RIGHTS. No Optionholder shall be deemed to be the holder of, or to have any of the rights of a holder with respect to, any shares subject to such Option unless and until such Optionholder has satisfied all requirements for exercise of the Option pursuant to its terms.

(b) NO SERVICE RIGHTS. Nothing in the Plan or any instrument executed or Option granted pursuant thereto shall confer upon any Optionholder any right to continue to serve the Company as a Non-Employee Director or shall affect the right of the Company or an Affiliate to terminate (i) the employment of an Employee with or without notice and with or without cause, (ii) the service of a Consultant pursuant to the terms of such Consultant's agreement with the Company or an Affiliate or (iii) the service of a Director pursuant to the Bylaws of the Company or an Affiliate, and any applicable provisions of the corporate law of the state in which the Company or the Affiliate is incorporated, as the case may be.

(c) INVESTMENT ASSURANCES. The Company may require an Optionholder, as a condition of exercising or acquiring stock under any Option, (i) to give written assurances satisfactory to the Company as to the Optionholder's knowledge and experience in financial and business matters and/or to employ a purchaser representative reasonably satisfactory to the Company who is knowledgeable and experienced in financial and business matters and that he or she is capable of evaluating, alone or together with the purchaser representative, the merits and risks of exercising the Option; and (ii) to give written assurances satisfactory to the Company stating that the Optionholder is acquiring the stock subject to the Option for the Optionholder's own account and not with any present intention of selling or otherwise distributing the stock. The foregoing requirements, and any assurances given pursuant to such requirements, shall be inoperative if (iii) the issuance of the shares upon the exercise or acquisition of

stock under the Option has been registered under a then currently effective registration statement under the Securities Act or (iv) as to any particular requirement, a determination is made by counsel for the Company that such requirement need not be met in the circumstances under the then applicable securities laws. The Company may, upon advice of counsel to the Company, place legends on stock certificates issued under the Plan as such counsel deems necessary or appropriate in order to comply with applicable securities laws, including, but not limited to, legends restricting the transfer of the stock.

(d) WITHHOLDING OBLIGATIONS. The Optionholder may satisfy any federal, state or local tax withholding obligation relating to the exercise or acquisition of stock under an Option by any of the following means (in addition to the Company's right to withhold from any compensation paid to the Optionholder by the Company) or by a combination of such means: (i) tendering a cash payment; (ii) authorizing the Company to withhold shares from the shares of the Common Stock otherwise issuable to the Optionholder as a result of the exercise or acquisition of stock under the Option, provided, however, that no shares of Common Stock are withheld with a value exceeding the minimum amount of tax required to be withheld by law; or (iii) delivering to the Company owned and unencumbered shares of the Common Stock.

11. ADJUSTMENTS UPON CHANGES IN STOCK.

(a) CAPITALIZATION ADJUSTMENTS. If any change is made in the stock subject to the Plan, or subject to any Option, without the receipt of consideration by the Company (through merger, consolidation, reorganization, recapitalization, reincorporation, stock dividend, dividend in property other than cash, stock split, liquidating dividend, combination of shares, exchange of shares, change in corporate structure or other transaction not involving the receipt of consideration by the Company), the Plan will be appropriately adjusted in the class(es) and number of securities subject to the Plan pursuant to subsection 4(a) and to be granted as nondiscretionary Options specified in Section 5, and the outstanding Options will be appropriately adjusted in the class(es) and number of securities and price per share of stock subject to such outstanding Options. The Board shall make such adjustments, and its determination shall be final, binding and conclusive. For clarity, the conversion of any convertible securities of the Company shall not be treated as a transaction "without receipt of consideration" by the Company.

(b) DISSOLUTION OR LIQUIDATION. In the event of a dissolution or liquidation of the Company, then all outstanding Options shall terminate immediately prior to such event.

(c) ASSET SALE, MERGER, CONSOLIDATION OR REVERSE MERGER.

(i) In the event of (i) a sale, lease or other disposition of all or substantially all of the assets of the Company, (ii) a merger or consolidation in which the Company is not the surviving corporation or (iii) a reverse merger in which the Company is the surviving corporation but the shares of Common Stock outstanding immediately preceding the merger are converted by virtue of the merger into other property, whether in the form of securities, cash or otherwise, then any surviving corporation or acquiring corporation shall assume any Options outstanding under the Plan or shall substitute similar Options (including an option to acquire the same consideration paid to the stockholders in the transaction described in this subsection 11(c) for those outstanding under the Plan).

(ii) In the event any surviving corporation or acquiring corporation refuses to assume such Options or to substitute similar Options for those outstanding under the Plan, then the vesting of such Options and the vesting of any shares of Common Stock acquired pursuant to such Options shall be accelerated in full, and the Options shall terminate if not exercised at or prior to such event.

(iii) In the event any surviving corporation or acquiring corporation assumes such Options or substitutes similar Options for those outstanding under the Plan but the Optionholder is not elected or appointed to the board of directors of the surviving corporation or acquiring corporation at the first meeting of such board of directors after such change in control event, then the vesting of such Options and the vesting of any shares of Common Stock acquired pursuant to such Options shall be accelerated by eighteen (18) months on the day after the first meeting of the board of directors of the surviving corporation or acquiring corporation.

(iv) In the event any surviving corporation or acquiring corporation assumes such Options or substitutes similar Options for those outstanding under the Plan and the Optionholder is elected or appointed to the board of directors of the surviving corporation or acquiring corporation at the first meeting of such board of directors after such change in control event, then the vesting of such Options and the vesting of any shares of Common Stock acquired pursuant to such Options shall not be accelerated.

12. AMENDMENT OF THE PLAN AND OPTIONS.

(a) AMENDMENT OF PLAN. The Board at any time, and from time to time, may amend the Plan. However, except as provided in Section 11 relating to adjustments upon changes in stock, no amendment shall be effective unless approved by the stockholders of the Company to the extent stockholder approval is necessary to satisfy the requirements of Rule 16b-3 or any Nasdaq or other securities exchange listing requirements.

(b) STOCKHOLDER APPROVAL. The Board may, in its sole discretion, submit any other amendment to the Plan for stockholder approval.

(c) NO IMPAIRMENT OF RIGHTS. Rights under any Option granted before amendment of the Plan shall not be impaired by any amendment of the Plan unless (i) the Company requests the consent of the Optionholder and (ii) the Optionholder consents in writing.

(d) AMENDMENT OF OPTIONS. The Board at any time, and from time to time, may amend the terms of any one or more Options; provided, however, that the rights under any Option shall not be impaired by any such amendment unless (i) the Company requests the consent of the Optionholder and (ii) the Optionholder consents in writing.

13. TERMINATION OR SUSPENSION OF THE PLAN.

(a) PLAN TERM. The Board may suspend or terminate the Plan at any time. Unless sooner terminated, the Plan shall terminate on the day before the tenth (10th) anniversary of the date the Plan is adopted by the Board or approved by the stockholders of the Company, whichever is earlier. No Options may be granted under the Plan while the Plan is suspended or after it is terminated.

(b) NO IMPAIRMENT OF RIGHTS. Suspension or termination of the Plan shall not impair rights and obligations under any Option granted while the Plan is in effect except with the written consent of the Optionholder.

14. EFFECTIVE DATE OF PLAN.

The Plan shall become effective as determined by the Board, but no Option shall be exercised unless and until the Plan has been approved by the stockholders of the Company, which approval shall be within twelve (12) months before or after the date the Plan is adopted by the Board.

15. CHOICE OF LAW.

All questions concerning the construction, validity and interpretation of this Plan shall be governed by the law of the State of Delaware, without regard to such state's conflict of laws rules.

LEXICON PHARMACEUTICALS, INC. 8800 TECHNOLOGY FOREST PLACE THE WOODLANDS, TX 77381-4287

VOTE BY INTERNET - www.proxyvote.com

Use the Internet to transmit your voting instructions and for electronic delivery of information up until 11:59 P.M. Eastern Time the day before the cut-off date or meeting date. Have your proxy card in hand when you access the web site and follow the instructions to obtain your records and to create an electronic voting instruction form.

ELECTRONIC DELIVERY OF FUTURE STOCKHOLDER COMMUNICATIONS

If you would like to reduce the costs incurred by Lexicon Pharmaceuticals, Inc. in mailing proxy materials, you can consent to receiving all future proxy statements, proxy cards and annual reports electronically via e-mail or the internet. To sign up for electronic delivery, please follow the instructions above to vote using the internet and, when prompted, indicate that you agree to receive or access stockholder communications electronically in future years.

VOTE BY PHONE - 1-800-690-6903

Use any touch-tone telephone to transmit your voting instructions up until 11:59 P.M. Eastern Time the day before the cut-off date or meeting date. Have your proxy card in hand when you call and then follow the instructions.

VOTE BY MAIL

Mark, sign and date your proxy card and return it in the postage-paid envelope we have provided or return it to Lexicon Pharmaceuticals, Inc. c/o Broadridge, 51 Mercedes Way, Edgewood, NY 11717.

TO VOTE, MARK BLOCKS BELOW IN BLUE OR LEXPH1 BLACK INK AS FOLLOWS

KEEP THIS PORTION FOR YOUR RECORDS DETACH AND RETURN THIS PORTION ONLY

THIS PROXY CARD IS VALID ONLY WHEN SIGNED AND DATED

LEXICON PHARMACEUTICALS, INC. For WithholdFor All To withhold authority to vote for All All Except any individual nominee(s), mark "For All Except" and write the number(s) of the nominee(s) on the line below.

THE BOARD OF o o o DIRECTORS RECOMMENDS A VOTE "FOR" ITEMS 1, 2, 3 AND 4

Vote on Directors

- 1. ELECTION OF
 - CLASS III

DIRECTORS

Nominees:

- Arthur T. 01) Sands, M.D., Ph.D.
- 02) Philippe J. Amouval
- Frank P. 03) Palantoni

Vote on Proposals

For Against Abstain

- 2. Approval of the Company's Equity Incentive Plan, amending and restating the 0 0 0 Company's existing 2000 Equity Incentive Plan 3. Approval of the Company's Non-Employee Directors' Stock Option Plan, amending 0 0 0 and restating the Company's existing 2000 Non-Employee Directors Stock Option Plan
- 4. Ratification and approval of the appointment of Ernst & Young LLP as the Company's independent auditors for the fiscal year ending December 31, 2009
- 5. In their discretion, upon such other matters that may properly come before the meeting or any adjournment or adjournments thereof.

The shares represented by this proxy, when properly executed, will be voted in the matter directed herein by the undersigned Stockholder(s). If no direction is made, this proxy will be voted FOR the election of the nominees listed in item 1 and for items 2, 3 and 4. If any other matters properly come before the meeting, or if cumulative voting is required, the person named in this proxy will vote in their discretion.

Please sign your name exactly as it appears hereon. When signing as attorney, executor, administrator, trustee or guardian, please add your title as such. When signing as joint tenants, all parties in the joint tenancy must sign. If a signer is a corporation, please sign in full corporate name by duly authorized officer.

Signature [PLEASE Date SIGN WITHIN BOX]

Signature (Joint Date Owners)

0 0 0 Important Notice Regarding the Availability of Proxy Materials for the Annual Meeting: The Notice and Proxy Statement and Annual Report are available at www.proxyvote.com.

LEXPH2

LEXICON PHARMACEUTICALS, INC.

THIS PROXY IS SOLICITED ON BEHALF OF THE BOARD OF DIRECTORS

ANNUAL MEETING OF STOCKHOLDERS April 23, 2009

The stockholder(s) hereby appoint(s) Arthur T. Sands, M.D., Ph.D. and Jeffrey L. Wade, or either of them, as proxies, each with the power to appoint his substitute, and hereby authorize(s) them to represent and to vote, as designated on the reverse side of this ballot, all of the shares of Common Stock of Lexicon Pharmaceuticals, Inc. that the stockholder(s) is/are entitled to vote at the Annual Meeting of Stockholders to be held at 1:30 p.m., local time, on April 23, 2009 at The Marriott Woodlands Waterway Hotel and Convention Center, 1601 Lake Robbins Drive, The Woodlands, Texas, and any adjournment or postponement thereof.

THIS PROXY, WHEN PROPERLY EXECUTED, WILL BE VOTED AS DIRECTED BY THE STOCKHOLDER(S). IF NO SUCH DIRECTIONS ARE MADE, THIS PROXY WILL BE VOTED FOR THE ELECTION OF THE NOMINEES LISTED ON THE REVERSE SIDE FOR THE BOARD OF DIRECTORS AND FOR EACH PROPOSAL.

PLEASE MARK, SIGN, DATE AND RETURN THIS PROXY CARD PROMPTLY USING THE ENCLOSED REPLY

CONTINUED AND TO BE SIGNED ON REVERSE SIDE